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(54) Title: NOVEL MOLECULES OF THE TANGO-77 RELATED PROTEIN FAMILY AND USES THEREOF				
(57) Abstract <p>Novel Tango-77 polypeptides, proteins, and nucleic acid molecules are disclosed. In addition to isolated, full-length Tango-77 proteins, the invention further provides isolated Tango-77 fusion proteins, antigenic peptides and anti-Tango-77 antibodies. The invention also provides Tango-77 nucleic acid molecules, recombinant expression vectors containing a nucleic acid molecule of the invention, host cells into which the expression vectors have been introduced and non-human transgenic animals in which a Tango-77 gene has been introduced or disrupted. Diagnostic, screening and therapeutic methods utilizing compositions of the invention are also provided.</p>				

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NOVEL MOLECULES OF THE TANGO-77 RELATED PROTEIN
FAMILY AND USES THEREOF

Background of the Invention

The polypeptide cytokine interleukin-1 (IL-1) is a critical mediator of inflammatory and overall immune response. To date, three members of the IL-1 family, IL-1 α , IL-1 β and IL-1ra (Interleukin-1 receptor antagonist) have been isolated and cloned. IL-1 α and IL-1 β are proinflammatory cytokines which elicit biological responses, whereas IL-1ra is an antagonist of IL-1 α and IL-1 β activity. Two distinct cell-surface receptors have been identified for these ligands, the type I IL-1 receptor (IL-1RtI) and type II IL-1 receptor (IL-1RtII). Recent results suggest that the IL-1RtI is the receptor responsible for transducing a signal and producing biological effects.

As mentioned above, IL-1 is a key mediator of the host inflammatory response. While inflammation is an important homeostatic mechanism, aberrant inflammation has the potential for inducing damage to the host. Elevated IL-1 levels are known to be associated with a number of diseases particularly autoimmune diseases and inflammatory disorders.

Since IL-1ra is a naturally occurring inhibitor of IL-1, IL-1ra can be used to limit the aberrant and potentially deleterious effects of IL-1. In experimental animals, pretreatment with IL-1ra has been shown to prevent death resulting from lipopolysaccharide-induced sepsis. The relative absence of IL-1ra has also been suggested to play a role in human inflammatory bowel disease.

Summary of the Invention

The present invention is based, at least in part, on the discovery of a gene encoding Tango-77, a secreted

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protein that is predicted to be a member of the cytokine superfamily. The Tango-77 cDNA described below (SEQ ID NO:1) has three possible open reading frames. The first potential open reading frame encompasses 534 nucleotides extending from nucleotide 356 to nucleotide 889 of SEQ ID NO:1 (SEQ ID NO:3) and encodes a 178 amino acid protein (SEQ ID NO:2). This protein may include a predicted signal sequence of about 63 amino acids (from about amino acid 1 to about amino acid 63 of SEQ ID NO:2 (SEQ ID NO:4) and a predicted mature protein of about 115 amino acids (from about amino acid 64 to amino acid 178 of SEQ ID NO:2 (SEQ ID NO:5)).

The second potential open reading frame encompasses 498 nucleotides extending from nucleotide 389 to nucleotide 889 of SEQ ID NO:1 (SEQ ID NO:6) and encodes a 167 amino acid protein (SEQ ID NO:7). This protein may include a predicted signal sequence of about 52 amino acids (from about amino acid 1 to about amino acid 52 of SEQ ID NO:7 (SEQ ID NO:8)) and a predicted mature protein of about 115 amino acids (from about amino acid 52 to amino acid 167 of SEQ ID NO:7 (SEQ ID NO:9)).

The third potential open reading frame encompasses 408 nucleotides extending from nucleotide 481 to nucleotide 889 of SEQ ID NO:1 (SEQ ID NO:10) and encodes a 136 amino acid protein (SEQ ID NO:11). This protein includes a predicted signal sequence of about 21 amino acids (from about amino acid 1 to about amino acid 21 of SEQ ID NO:11 (SEQ ID NO:12)) and a predicted mature protein of about 115 amino acids (from about amino acid 22 to amino acid 136 of SEQ ID NO:11 (SEQ ID NO:13)).

As used herein, the terms "Tango-77", "Tango-77 protein", "Tango-77 polypeptide" and the like, can refer and polypeptide produced by the cDNA of SEQ ID NO:1 including any and all of the Tango-77 gene products described above.

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Tango-77 is expected to inhibit inflammation and play a functional role similar to that of secreted IL-1ra. For example, it is expected that Tango-77 may bind to the IL-1 receptor, thus blocking receptor activation by inhibiting the binding of IL-1 α and IL-1 β to the receptor. Alternatively, Tango-77 may inhibit inflammation through another pathway, for example, by binding to a novel receptor. Accordingly, Tango-77 may be useful as a modulating agent in regulating a variety of cellular processes including acute and chronic inflammation, e.g., asthma, chronic myelogenous leukemia, rheumatoid arthritis, psoriasis and inflammatory bowel disease.

In one aspect, the invention provides isolated nucleic acid molecules encoding Tango-77 or biologically active portions thereof, as well as nucleic acid fragments suitable as primers or hybridization probes for the detection of Tango-77.

The invention encompasses methods of diagnosing and treating patients who are suffering from a disorder associated with an abnormal level (undesirably high or undesirably low) of inflammation, abnormal activity of the IL-1 receptor complex, or abnormal activity of IL-1, by administering a compound that modulates the expression of Tango-77 (at the DNA, mRNA or protein level, e.g., by altering mRNA splicing) or by altering the activity of Tango-77. Examples of such compounds include small molecules, antisense nucleic acid molecules, ribozymes, and polypeptides.

The invention features a nucleic acid molecule which is at least 45% (e.g., 55%, 65%, 75%, 85%, 95%, or 98%) identical to the nucleotide sequence shown in SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the nucleotide sequence of the cDNA insert of the plasmid

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deposited with ATCC as Accession Number (the "cDNA of ATCC 98807"), or a complement thereof.

The invention features a nucleic acid molecule which includes a fragment of at least 100 (e.g., 250,
5 325, 350, 375, 400, 425, 450, 500, 550, 600, 650, 700, 800, 900, or 989) nucleotides of the nucleotide sequence shown in SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the nucleotide sequence of the cDNA ATCC 98807, or a complement thereof.

10 The invention also features a nucleic acid molecule which includes a nucleotide sequence encoding a protein having an amino acid sequence that is at least 45% (55%, 65%, 75%, 85%, 95%, or 98%) identical to the amino acid sequence of SEQ ID NO:2, SEQ ID NO:5, SEQ ID
15 NO:7, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:13, or the amino acid sequence encoded by the cDNA of ATCC 98807.

In a preferred embodiment, a Tango-77 nucleic acid molecule has the nucleotide sequence shown in SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10 or the
20 nucleotide sequence of the cDNA of ATCC 98807.

Also within the invention is a nucleic acid molecule which encodes a fragment of a polypeptide having the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID
25 NO:11, SEQ ID NO:12, SEQ ID NO:13, wherein the fragment includes at least 15 (e.g., 25, 30, 50, 100, 150, or 178) contiguous amino acids of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or the polypeptide
30 encoded by the cDNA of ATCC Accession Number 98807.

The invention includes a nucleic acid molecule which encodes a naturally occurring allelic variant of a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8,
35 SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or

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an amino acid sequence encoded by the cDNA of ATCC Accession Number 98807, wherein the nucleic acid molecule hybridizes to a nucleic acid molecule comprising SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, or a complement thereof under stringent conditions.

Also within the invention are: an isolated Tango-77 protein having an amino acid sequence that is at least about 45%, preferably 65%, 75%, 85%, 95%, or 98% identical to the amino acid sequence of SEQ ID NO:5, SEQ ID NO:9 or SEQ ID NO:13 (mature human Tango-77), or the amino acid sequence of SEQ ID NO:2, SEQ ID NO:7 or SEQ ID NO:11 (immature human Tango-77).

Also within the invention are: an isolated Tango-77 protein which is encoded by a nucleic acid molecule having a nucleotide sequence that is at least about 65%, preferably 75%, 85%, or 95% identical to SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10 or the cDNA of ATCC 98807; and an isolated Tango-77 protein which is encoded by a nucleic acid molecule having a nucleotide sequence which hybridizes under stringent hybridization conditions to a nucleic acid molecule having the nucleotide sequence of SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the non-coding strand of the cDNA of ATCC 98807, or the complement thereof.

Also within the invention is a polypeptide which is a naturally occurring allelic variant of a polypeptide that includes the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807, wherein the polypeptide is encoded by a nucleic acid molecule which hybridizes to a nucleic acid molecule comprising SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID

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NO:10 or the complement thereof under stringent conditions.

Another embodiment of the invention features Tango-77 nucleic acid molecules which specifically detect
5 Tango-77 nucleic acid molecules relative to nucleic acid molecules encoding other members of the cytokine superfamily. For example, in one embodiment, a Tango-77 nucleic acid molecule hybridizes under stringent conditions to a nucleic acid molecule comprising the
10 nucleotide sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the cDNA of ATCC 98807, or a complement thereof. In another embodiment, the Tango-77 nucleic acid molecule is at least 300 (325, 350, 375, 400, 425, 450, 500, 550, 600, 650, 700, 800, 900, or 989)
15 nucleotides in length and hybridizes under stringent conditions to a nucleic acid molecule comprising the nucleotide sequence shown in SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the cDNA of ATCC 98807, or a complement thereof. In yet another embodiment, the
20 invention provides an isolated nucleic acid molecule which is antisense to the coding strand of a Tango-77 nucleic acid.

Another aspect of the invention provides a vector, e.g., a recombinant expression vector, comprising a
25 Tango-77 nucleic acid molecule of the invention. In another embodiment, the invention provides a host cell containing such a vector. The invention also provides a method for producing Tango-77 protein by culturing, in a suitable medium, a host cell of the invention containing
30 a recombinant expression vector such that a Tango-77 protein is produced.

Another aspect of this invention features isolated or recombinant Tango-77 proteins and polypeptides. Preferred Tango-77 proteins and polypeptides possess at
35 least one biological activity possessed by naturally

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occurring human Tango-77, e.g., (i) the ability to interact with proteins in the Tango-77 signalling pathway (ii) the ability to interact with a Tango-77 ligand or receptor; or (iii) the ability to interact with an intracellular target protein, (iv) the ability to
5 interact with a protein involved in inflammation and (v) the ability to bind the IL-1 receptor. Other activities include the induction and suppression of polypeptide interleukins, cytokines and growth factors.

10 The Tango-77 proteins of the present invention, or biologically active portions thereof, can be operably linked to a non-Tango-77 polypeptide (e.g., heterologous amino acid sequences) to form Tango-77 fusion proteins. The invention further features antibodies that
15 specifically bind Tango-77 proteins, such as monoclonal or polyclonal antibodies. In addition, the Tango-77 proteins or biologically active portions thereof can be incorporated into pharmaceutical compositions, which optionally include pharmaceutically acceptable carriers.

20 In another aspect, the present invention provides a method for detecting the presence of Tango-77 activity or expression in a biological sample by contacting the biological sample with an agent capable of detecting an indicator of Tango-77 activity or expression such that
25 the presence of Tango-77 activity or expression is detected in the biological sample.

In another aspect, the invention provides a method for modulating Tango-77 activity comprising contacting a cell with an agent that modulates (inhibits or
30 stimulates)

Tango-77 activity or expression such that Tango-77 activity or expression in the cell is modulated. In one embodiment, the agent is an antibody that specifically binds to Tango-77 protein. In another embodiment, the

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agent modulates expression of Tango-77 by modulating transcription of a Tango-77 gene, splicing of a Tango-77 mRNA, or translation of a Tango-77 mRNA. In yet another embodiment, the agent is a nucleic acid molecule having a nucleotide sequence that is antisense to the coding strand of the Tango-77 mRNA or the Tango-77 gene.

In one embodiment, the methods of the present invention are used to treat a subject having a disorder characterized by aberrant Tango-77 protein activity or nucleic acid expression by administering an agent which is a Tango-77 modulator to the subject. In one embodiment, the Tango-77 modulator is a Tango-77 protein. In another embodiment, the Tango-77 modulator is a Tango-77 nucleic acid molecule. In other embodiments, the Tango-77 modulator is a peptide, peptidomimetic, or other small molecule. In a preferred embodiment, the disorder characterized by aberrant Tango-77 protein or nucleic acid expression can include chronic and acute inflammation.

The present invention also provides a diagnostic assay for identifying the presence or absence of a genetic lesion or mutation characterized by at least one of: (i) aberrant modification or mutation of a gene encoding a Tango-77 protein; (ii) mis-regulation of a gene encoding a Tango-77 protein; and (iii) aberrant post-translational modification of a Tango-77 protein, wherein a wild-type form of the gene encodes a protein with a Tango-77 activity.

In another aspect, the invention provides a method for identifying a compound that binds to or modulates the activity of a Tango-77 protein. In general, such methods entail measuring a biological activity of a Tango-77 protein in the presence and absence of a test compound and identifying those

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compounds which alter the activity of the Tango-77 protein.

The invention also features methods for identifying a compound which modulates the expression of Tango-77 by measuring the expression of Tango-77 in the presence and absence of a compound.

Other features and advantages of the invention will be apparent from the following detailed description and claims.

Brief Description of the Drawings

Figure 1 depicts the cDNA sequence (SEQ ID NO:1) of Tango-77. The Tango-77 cDNA has three possible open reading frames which encode the amino acid sequence (SEQ ID NO:2, SEQ ID NO:7 and SEQ ID NO:11) of human Tango-77. The three potential open reading frames of SEQ ID NO:1 extend from: (1) nucleotide 356 to nucleotide 889 (SEQ ID NO:3); (2) nucleotide 389 to nucleotide 889 (SEQ ID NO:6); and (3) nucleotide 481 to nucleotide 889 (SEQ ID NO:10).

Figure 2 depicts an alignment of an amino acid sequence of Tango-77 (T77; SEQ ID NO:2) with IL-1RA (SEQ ID NO:14), and IL-1 β (SEQ ID NO:15).

Figure 3 depicts the genomic sequence of BAC1 (SEQ ID NO:16).

Figure 4 depicts the genomic sequence of BAC2 (SEQ ID NO:17).

Figure 5 depicts an amino acid sequence of an alternatively spliced form of Tango-77 (SEQ ID NO:2) as predicted by Procrustes (T77-procrustes; SEQ ID NO:18).

Figure 6 depicts an alignment of an amino acid sequence of an alternatively spliced form of Tango-77 (T77-procrustes; SEQ ID NO:18) with Tango-77 (SEQ ID NO:2).

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Figure 7 depicts an alignment of an amino acid sequence of an alternatively spliced form of Tango-77 (T77-procrustes; SEQ ID NO:18) with IL-1ra (SEQ ID NO:14), and IL-1 β (SEQ ID NO:15).

5 Detailed Description of the Invention

The present invention is based on the discovery of a cDNA molecule encoding human Tango-77, a member of the cytokine superfamily. The cDNA molecule encoding human Tango-77 has three possible open reading frames. The
10 three possible nucleotide open reading frames for human Tango-77 protein are shown in Figure 1 (SEQ ID NO:3, SEQ ID NO:6 and SEQ ID NO:10). The predicted amino acid sequence for the three possible Tango-77 immature proteins are also shown in
15 Figure 1 (SEQ ID NO:2, SEQ ID NO:7 or SEQ ID NO:11) and three possible mature proteins are also shown in Figure 1 (SEQ ID NO:5, SEQ ID NO:9 and SEQ ID NO:13).

The Tango-77 cDNA of Figure 1 (SEQ ID NO:1), which is approximately 989 nucleotides long including
20 untranslated regions, encodes a protein amino acid having a molecular weight of approximately 19 kDa, 18 kDa, or 14.9 kDa (excluding post-translational modifications) and the possible mature form of the protein has a molecular weight of 13 kDa. A plasmid containing a cDNA encoding
25 human Tango-77 (with the cDNA insert name of Of fthx077) was deposited with American Type Culture Collection (ATCC), 10801 University Boulevard, Manassas, Virginia 20110-2209 on July 2, 1998 and assigned Accession Number 98807. This deposit will be maintained under the terms
30 of the Budapest Treaty on the International Recognition of the Deposit of Microorganisms for the Purposes of Patent Procedure. This deposit was made merely as a convenience for those of skill in the art and is not an

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admission that a deposit is required under 35 U.S.C. §112.

Human Tango-77 is one member of a family of molecules (the "Tango-77 family") having certain
5 conserved structural and functional features. The term "family," when referring to the protein and nucleic acid molecules of the invention, is intended to mean two or more proteins or nucleic acid molecules having a common structural domain and having sufficient amino acid or
10 nucleotide sequence identity as defined herein. Such family members can be naturally occurring and can be from either the same or different species. For example, a family can contain a first protein of human origin and a homologue of that protein of murine origin, as well as a
15 second, distinct protein of human origin and a murine homologue of that protein. Members of a family may also have common functional characteristics.

As used interchangeably herein a "Tango-77 activity", "biological activity of Tango-77" or
20 "functional activity of Tango-77", refers to an activity exerted by a Tango-77 protein, polypeptide or nucleic acid molecule on a Tango-77 responsive cell as determined *in vivo*, or *in vitro*, according to standard techniques. A Tango-77 activity can be a direct activity, such as an
25 association with a second protein, or an indirect activity, such as a cellular signaling activity mediated by interaction of the Tango-77 protein with a second protein. In a preferred embodiment, a Tango-77 activity includes at least one or more of the following
30 activities: (i) the ability to interact with proteins in the Tango-77 signalling pathway (ii) the ability to interact with a Tango-77 ligand or receptor; or (iii) the ability to interact with an intracellular target protein, (iv) the ability to interact with a protein involved in

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inflammation, and (v) the ability to bind the IL-1 receptor.

Accordingly, another embodiment of the invention features isolated Tango-77 proteins and polypeptides
5 having a Tango-77 activity.

Yet another embodiment of the invention features Tango-77 molecules which contain a signal sequence. Generally, a signal sequence (or signal peptide) is a peptide containing about 21 to 63 amino acids which
10 occurs at the extreme N-terminal end of a secretory protein. The native Tango-77 signal sequence (SEQ ID NO:4, SEQ ID NO:8, or SEQ ID NO:12) can be removed and replaced with a signal sequence from another protein. In certain host cells (e.g., mammalian host cells),
15 expression and/or secretion of Tango-77 can be increased through use of a heterologous signal sequence. For example, the gp67 secretory sequence of the baculovirus envelope protein can be used as a heterologous signal sequence. Alternatively, the native Tango-77 signal
20 sequence can itself be used as a heterologous signal sequence in expression systems, e.g., to facilitate the secretion of a protein of interest.

Various aspects of the invention are described in further detail in the following subsections.

25 I. Isolated Nucleic Acid Molecules

One aspect of the invention pertains to isolated nucleic acid molecules that encode Tango-77 proteins or biologically active portions thereof, as well as nucleic acid molecules sufficient for use as hybridization probes
30 to identify Tango-77-encoding nucleic acids (e.g., Tango-77 mRNA) and fragments for use as PCR primers for the amplification or mutation of Tango-77 nucleic acid molecules. As used herein, the term "nucleic acid molecule" is intended to include DNA molecules (e.g.,

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cDNA or genomic DNA) and RNA molecules (e.g., mRNA) and analogs of the DNA or RNA generated using nucleotide analogs. The nucleic acid molecule can be single-stranded or double-stranded, but preferably is double-stranded DNA.

An "isolated" nucleic acid molecule is one which is separated from other nucleic acid molecules which are present in the natural source of the nucleic acid. Preferably, an "isolated" nucleic acid is free of sequences (preferably protein encoding sequences) which naturally flank the nucleic acid (i.e., sequences located at the 5' and 3' ends of the nucleic acid) in the genomic DNA of the organism from which the nucleic acid is derived. For example, in various embodiments, the isolated Tango-77 nucleic acid molecule can contain less than about 5 kb, 4 kb, 3 kb, 2 kb, 1 kb, 0.5 kb or 0.1 kb of nucleotide sequences which naturally flank the nucleic acid molecule in genomic DNA of the cell from which the nucleic acid is derived. Moreover, an "isolated" nucleic acid molecule, such as a cDNA molecule, can be substantially free of other cellular material, or culture medium when produced by recombinant techniques, or substantially free of chemical precursors or other chemicals when chemically synthesized.

A nucleic acid molecule of the present invention, e.g., a nucleic acid molecule having the nucleotide sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the cDNA of ATCC 98807, or a complement of any of these nucleotide sequences, can be isolated using standard molecular biology techniques and the sequence information provided herein. Using all or a portion of the nucleic acid sequences of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the cDNA of ATCC 98807, or the complement thereof as a hybridization probe, Tango-77 nucleic acid molecules can be isolated using standard

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hybridization and cloning techniques (e.g., as described in Sambrook et al., eds., *Molecular Cloning: A Laboratory Manual*, 2nd ed., Cold Spring Harbor Laboratory, Cold Spring Harbor Laboratory Press, Cold Spring Harbor, NY, 1989).

A nucleic acid of the invention can be amplified using cDNA, mRNA or genomic DNA as a template and appropriate oligonucleotide primers according to standard PCR amplification techniques. The nucleic acid so amplified can be cloned into an appropriate vector and characterized by DNA sequence analysis. Furthermore, oligonucleotides corresponding to Tango-77 nucleotide sequences can be prepared by standard synthetic techniques, e.g., using an automated DNA synthesizer.

In another preferred embodiment, an isolated nucleic acid molecule of the invention comprises a nucleic acid molecule which is a complement of the nucleotide sequence shown in SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10 the cDNA of ATCC 98807, or a portion thereof. A nucleic acid molecule which is complementary to a given nucleotide sequence is one which is sufficiently complementary to the given nucleotide sequence that it can hybridize to the given nucleotide sequence thereby forming a stable duplex.

Moreover, the nucleic acid molecule of the invention can comprise only a portion of a nucleic acid sequence encoding Tango-77, for example, a fragment which can be used as a probe or primer or a fragment encoding a biologically active portion of Tango-77. The nucleotide sequence determined from the cloning of the human Tango-77 gene allows for the generation of probes and primers designed for use in identifying and/or cloning Tango-77 homologues in other cell types, e.g., from other tissues, as well as Tango-77 homologues from other mammals. The probe/primer typically comprises

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substantially purified oligonucleotide. The oligonucleotide typically comprises a region of nucleotide sequence that hybridizes under stringent conditions to at least about 12, preferably about 25,
5 more preferably about 50, 75, 100, 125, 150, 175, 200, 250, 300, 350 or 400 consecutive nucleotides of the sense or anti-sense sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, or the cDNA of ATCC 98807. Alternatively, the oligonucleotide can typically comprise
10 a region of nucleotide sequence that hybridizes under stringent conditions to at least about 12, preferably about 25, more preferably about 50, 75, 100, 125, 150, 175, 200, 250, 300, 350 or 400 consecutive nucleotides of the sense or anti-sense sequence of a naturally occurring
15 mutant of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, or the cDNA of ATCC 98807.

Probes based on the human Tango-77 nucleotide sequence can be used to detect transcripts or genomic sequences encoding the same or identical proteins. The
20 probe comprises a label group attached thereto, e.g., a radioisotope, a fluorescent compound, an enzyme, or an enzyme co-factor. Such probes can be used as a part of a diagnostic test kit for identifying cells or tissues which mis-express a Tango-77 protein, such as by
25 measuring a level of a Tango-77-encoding nucleic acid in a sample of cells from a subject, e.g., detecting Tango-77 mRNA levels or determining whether a genomic Tango-77 gene has been mutated or deleted.

A nucleic acid fragment encoding a "biologically
30 active portion of Tango-77" can be prepared by isolating a portion of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10 or the nucleotide sequence of the cDNA of ATCC 98807 which encodes a polypeptide having a Tango-77 biological activity, expressing the encoded portion of
35 Tango-77 protein (e.g., by recombinant expression in

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vitro) and assessing the activity of the encoded portion of Tango-77.

The invention further encompasses nucleic acid molecules that differ from the nucleotide sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, or the cDNA of ATCC 98807 due to degeneracy of the genetic code and thus encode the same Tango-77 protein as that encoded by the nucleotide sequence shown in SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, or the cDNA of ATCC 98807.

In addition to the human Tango-77 nucleotide sequence shown in SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, or the cDNA of ATCC 98807, it will be appreciated by those skilled in the art that DNA sequence polymorphisms that lead to changes in the amino acid sequences of Tango-77 may exist within a population (e.g., the human population). Such genetic polymorphism in the Tango-77 gene may exist among individuals within a population due to natural allelic variation. An allele is one of a group of genes which occur alternatively at a given genetic locus. As used herein, the term "allelic variant" refers to a nucleotide sequence which occurs at a Tango-77 locus or to a polypeptide encoded by the nucleotide sequence. As used herein, the terms "gene" and "recombinant gene" refer to nucleic acid molecules comprising an open reading frame encoding a Tango-77 protein, preferably a mammalian Tango-77 protein. Such natural allelic variations can typically result in 1-5% variance in the nucleotide sequence of the Tango-77 gene. Alternative alleles can be identified by sequencing the gene of interest in a number of different individuals. This can be readily carried out by using hybridization probes to identify the same genetic locus in a variety of individuals. Any and all such nucleotide variations and resulting amino acid polymorphisms or variations in

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Tango-77 that are the result of natural allelic variation and that do not alter the functional activity of Tango-77 are intended to be within the scope of the invention.

Moreover, nucleic acid molecules encoding Tango-77
5 proteins from other species (Tango-77 homologues), which have a nucleotide sequence which differs from that of a human Tango-77, are intended to be within the scope of the invention. Nucleic acid molecules corresponding to natural allelic variants and homologues of the Tango-77
10 cDNA of the invention can be isolated based on their identity to the human Tango-77 nucleic acids disclosed herein using the human cDNAs, or a portion thereof, as a hybridization probe according to standard hybridization techniques under stringent hybridization conditions.

15 Accordingly, in another embodiment, an isolated nucleic acid molecule of the invention is at least 300 (325, 350, 375, 400, 425, 450, 500, 550, 600, 650, 700, 800, or 989) nucleotides in length and hybridizes under stringent conditions to the nucleic acid molecule
20 comprising the nucleotide sequence, preferably the coding sequence, of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, or the cDNA of ATCC 98807.

As used herein, the term "hybridizes under stringent conditions" is intended to describe conditions
25 for hybridization and washing under which nucleotide sequences at least 60% (65%, 70%, preferably 75%) identical to each other typically remain hybridized to each other. Such stringent conditions are known to those skilled in the art and can be found in *Current Protocols*
30 in *Molecular Biology*, John Wiley & Sons, N.Y. (1989), 6.3.1-6.3.6. A preferred, non-limiting example of stringent hybridization conditions are hybridization in 6X sodium chloride/sodium citrate (SSC) at about 45°C, followed by one or more washes in 0.2X SSC, 0.1% SDS at
35 50-65°C. Preferably, an isolated nucleic acid molecule

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of the invention that hybridizes under stringent conditions to the sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the cDNA of ATCC 98807, or the complement thereof, corresponds to a naturally-occurring
5 nucleic acid molecule. As used herein, a "naturally-occurring" nucleic acid molecule refers to an RNA or DNA molecule having a nucleotide sequence that occurs in nature (e.g., encodes a natural protein).

In addition to naturally-occurring allelic
10 variants of the Tango-77 sequence that may exist in the population, the skilled artisan will further appreciate that changes can be introduced by mutation into the nucleotide sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10 or the cDNA of ATCC 98807, thereby
15 leading to changes in the amino acid sequence of the encoded Tango-77 protein, without altering the biological activity of the Tango-77 protein. Amino acid residues that are not conserved or only semiconserved among Tango-77 of various species may be non-essential for activity
20 and thus would likely be targets for alteration. Alternatively, one can make nucleotide substitutions leading to amino acid substitutions at "non-essential" amino acid residues. A "non-essential" amino acid residue is a residue that can be altered from the wild-
25 type sequence of Tango-77 (e.g., the sequence of SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11 or SEQ ID NO:13) without altering the biological activity, whereas an "essential" amino acid residue is required for biological activity. For example, amino
30 acid residues that are conserved among the Tango-77 proteins of various species may be essential for activity and thus would not likely be targets for alteration, unless one wishes to reduce or alter Tango-77 activity.

Accordingly, another aspect of the invention
35 pertains to nucleic acid molecules encoding Tango-77

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proteins that contain changes in amino acid residues that are not essential for activity. Such Tango-77 proteins differ in amino acid sequence from SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13 yet retain biological activity. In one embodiment, the isolated nucleic acid molecule includes a nucleotide sequence encoding a protein that includes an amino acid sequence that is at least about 45% identical, 65%, 75%, 85%, 95%, or 98% identical to the amino acid sequence of SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13.

An isolated nucleic acid molecule encoding a Tango-77 protein having a sequence which differs from that of SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13 can be created by introducing one or more nucleotide substitutions, additions or deletions into the nucleotide sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, or the cDNA of ATCC 98807 such that one or more amino acid substitutions, additions or deletions are introduced into the encoded protein. Mutations can be introduced by standard techniques, such as site-directed mutagenesis and PCR-mediated mutagenesis. Preferably, conservative amino acid substitutions are made at one or more predicted non-essential amino acid residues. A "conservative amino acid substitution" is one in which the amino acid residue is replaced with an amino acid residue having a similar side chain. Families of amino acid residues having similar side chains have been defined in the art. These families include amino acids with basic side chains (e.g., lysine, arginine, histidine), acidic side chains (e.g., aspartic acid, glutamic acid), uncharged polar side chains (e.g., glycine, asparagine, glutamine, serine, threonine, tyrosine, cysteine), nonpolar side chains (e.g., alanine,

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valine, leucine, isoleucine, proline, phenylalanine, methionine, tryptophan), beta-branched side chains (e.g., threonine, valine, isoleucine) and aromatic side chains (e.g., tyrosine, phenylalanine, tryptophan, histidine).

5 Thus, a predicted nonessential amino acid residue in Tango-77 is preferably replaced with another amino acid residue from the same side chain family. Alternatively, mutations can be introduced randomly along all or part of a Tango-77 coding sequence, such as by saturation
10 mutagenesis, and the resultant mutants can be screened for Tango-77 biological activity to identify mutants that retain activity. Following mutagenesis, the encoded protein can be expressed recombinantly and the activity of the protein can be determined.

15 In a preferred embodiment, a mutant Tango-77 protein can be assayed for: (1) the ability to form protein:protein interactions with proteins in the Tango-77 signalling pathway; (2) the ability to bind a Tango-77 ligand or receptor; or (3) the ability to bind
20 to an intracellular target protein or (4) the ability to interact with a protein involved in inflammation or (5) the ability to bind the IL-1 receptor. In yet another preferred embodiment, a mutant Tango-77 can be assayed for the ability to modulate inflammation, asthma,
25 autoimmune diseases, and sepsis.

The present invention encompasses antisense nucleic acid molecules, i.e., molecules which are complementary to a sense nucleic acid encoding a protein, e.g., complementary to the coding strand of a double-
30 stranded cDNA molecule or complementary to an mRNA sequence. Accordingly, an antisense nucleic acid can hydrogen bond to a sense nucleic acid. The antisense nucleic acid can be complementary to an entire Tango-77 coding strand, or to only a portion thereof, e.g., all or
35 part of the protein coding region (or open reading

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frame). An antisense nucleic acid molecule can be antisense to a noncoding region of the coding strand of a nucleotide sequence encoding Tango-77. The noncoding regions ("5' and 3' untranslated regions") are the 5' and 3' sequences which flank the coding region and are not translated into amino acids.

Given the coding strand sequences encoding Tango-77 disclosed herein (e.g., SEQ ID NO:3, SEQ ID NO:5, or SEQ ID NO:8), antisense nucleic acids of the invention can be designed according to the rules of Watson and Crick base pairing. The antisense nucleic acid molecule can be complementary to the entire coding region of Tango-77 mRNA, but more preferably is an oligonucleotide which is antisense to only a portion of the coding or noncoding region of Tango-77 mRNA. For example, the antisense oligonucleotide can be complementary to the region surrounding the translation start site of Tango-77 mRNA, e.g., an oligonucleotide having the sequence 5'-TGCAACTTTTACAGGAAACAC-3' (SEQ ID NO:19) or 5'-CCTCACTTTTACCCGAGACTC-3' (SEQ ID NO:20) or 5'-GACGGGTGGTACTTAAACAA-3' (SEQ ID NO:21). An antisense oligonucleotide can be, for example, about 5, 10, 15, 20, 25, 30, 35, 40, 45 or 50 nucleotides in length. An antisense nucleic acid of the invention can be constructed using chemical synthesis and enzymatic ligation reactions using procedures known in the art. For example, an antisense nucleic acid (e.g., an antisense oligonucleotide) can be chemically synthesized using naturally occurring nucleotides or variously modified nucleotides designed to increase the biological stability of the molecules or to increase the physical stability of the duplex formed between the antisense and sense nucleic acids, e.g., phosphorothioate derivatives and acridine substituted nucleotides can be used. Examples of modified nucleotides which can be used to

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generate the antisense nucleic acid include 5-fluorouracil, 5-bromouracil, 5-chlorouracil, 5-iodouracil, hypoxanthine, xanthine, 4-acetylcytosine, 5-(carboxyhydroxymethyl) uracil, 5-carboxymethylaminomethyl-2-thiouridine, 5-carboxymethylaminomethyluracil, dihydrouracil, beta-D-galactosylqueosine, inosine, N6-isopentenyladenine, 1-methylguanine, 1-methylinosine, 2,2-dimethylguanine, 2-methyladenine, 2-methylguanine, 3-methylcytosine, 5-methylcytosine, N6-adenine, 7-methylguanine, 5-methylaminomethyluracil, 5-methoxyaminomethyl-2-thiouracil, beta-D-mannosylqueosine, 5'-methoxycarboxymethyluracil, 5-methoxyuracil, 2-methylthio-N6-isopentenyladenine, uracil-5-oxyacetic acid (v), wybutoxosine, pseudouracil, queosine, 2-thiocytosine, 5-methyl-2-thiouracil, 2-thiouracil, 4-thiouracil, 5-methyluracil, uracil-5-oxyacetic acid methylester, uracil-5-oxyacetic acid (v), 5-methyl-2-thiouracil, 3-(3-amino-3-N-2-carboxypropyl) uracil (acp3)w, and 2,6-diaminopurine. Alternatively, the antisense nucleic acid can be produced biologically using an expression vector into which a nucleic acid has been subcloned in an antisense orientation (i.e., RNA transcribed from the inserted nucleic acid will be of an antisense orientation to a target nucleic acid of interest, described further in the following subsection).

The antisense nucleic acid molecules of the invention are typically administered to a subject or generated *in situ* such that they hybridize with or bind to cellular mRNA and/or genomic DNA encoding a Tango-77 protein to thereby inhibit expression of the protein, e.g., by inhibiting transcription and/or translation. The hybridization can be by conventional nucleotide complementarity to form a stable duplex, or, for example, in the case of an antisense nucleic acid molecule which

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binds to DNA duplexes, through specific interactions in the major groove of the double helix. An example of a route of administration of antisense nucleic acid molecules of the invention includes direct injection at a tissue site. Alternatively, antisense nucleic acid molecules can be modified to target selected cells and then administered systemically. For example, for systemic administration, antisense molecules can be modified such that they specifically bind to receptors or antigens expressed on a selected cell surface, e.g., by linking the antisense nucleic acid molecules to peptides or antibodies which bind to cell surface receptors or antigens. The antisense nucleic acid molecules can also be delivered to cells using the vectors described herein.

To achieve sufficient intracellular concentrations of the antisense molecules, vector constructs in which the antisense nucleic acid molecule is placed under the control of a strong pol II or pol III promoter are preferred.

An antisense nucleic acid molecule of the invention can be an α -anomeric nucleic acid molecule. An α -anomeric nucleic acid molecule forms specific double-stranded hybrids with complementary RNA in which, contrary to the usual β -units, the strands run parallel to each other (Gaultier et al. (1987) *Nucleic Acids Res.* 15:6625-6641). The antisense nucleic acid molecule can also comprise a 2'-o-methylribonucleotide (Inoue et al. (1987) *Nucleic Acids Res.* 15:6131-6148) or a chimeric RNA-DNA analogue (Inoue et al. (1987) *FEBS Lett.* 215:327-330).

The invention also encompasses ribozymes. Ribozymes are catalytic RNA molecules with ribonuclease activity which are capable of cleaving a single-stranded nucleic acid, such as an mRNA, to which they have a complementary region. Thus, ribozymes (e.g., hammerhead

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ribozymes (described in Haselhoff and Gerlach (1988) *Nature* 334:585-591)) can be used to catalytically cleave Tango-77 mRNA transcripts to thereby inhibit translation of Tango-77 mRNA. A ribozyme having specificity for a Tango-77-encoding nucleic acid can be designed based upon the nucleotide sequence of a Tango-77 cDNA disclosed herein (e.g., SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10). For example, a derivative of a *Tetrahymena* L-19 IVS RNA can be constructed in which the nucleotide sequence of the active site is complementary to the nucleotide sequence to be cleaved in a Tango-77-encoding mRNA. See, e.g., Cech et al. U.S. Patent No. 4,987,071; and Cech et al. U.S. Patent No. 5,116,742. Alternatively, Tango-77 mRNA can be used to select a catalytic RNA having a specific ribonuclease activity from a pool of RNA molecules. See, e.g., Bartel and Szostak (1993) *Science* 261:1411-1418.

The invention also encompasses nucleic acid molecules which form triple helical structures. For example, Tango-77 gene expression can be inhibited by targeting nucleotide sequences complementary to the regulatory region of the Tango-77 (e.g., the Tango-77 promoter and/or enhancers) to form triple helical structures that prevent transcription of the Tango-77 gene in target cells. See generally, Helene (1991) *Anticancer Drug Des.* 6(6):569-84; Helene (1992) *Ann. N.Y. Acad. Sci.* 660:27-36; and Maher (1992) *Bioassays* 14(12):807-15.

In preferred embodiments, the nucleic acid molecules of the invention can be modified at the base moiety, sugar moiety or phosphate backbone to improve, e.g., the stability, hybridization, or solubility of the molecule. For example, the deoxyribose phosphate backbone of the nucleic acids can be modified to generate peptide nucleic acids (see Hyrup et al. (1996) *Bioorganic*

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& *Medicinal Chemistry* 4(1): 5-23). As used herein, the terms "peptide nucleic acids" or "PNAs" refer to nucleic acid mimics, e.g., DNA mimics, in which the deoxyribose phosphate backbone is replaced by a pseudopeptide backbone and only the four natural nucleobases are retained. The neutral backbone of PNAs has been shown to allow for specific hybridization to DNA and RNA under conditions of low ionic strength. The synthesis of PNA oligomers can be performed using standard solid phase peptide synthesis protocols as described in Hyrup et al. (1996) *supra*; Perry-O'Keefe et al. (1996) *Proc. Natl. Acad. Sci. USA* 93: 14670-675.

PNAs of Tango-77 can be used in therapeutic and diagnostic applications. For example, PNAs can be used as antisense or antigene agents for sequence-specific modulation of gene expression by, e.g., inducing transcription or translation arrest or inhibiting replication. PNAs of Tango-77 can also be used, e.g., in the analysis of single base pair mutations in a gene by, e.g., PNA directed PCR clamping; as artificial restriction enzymes when used in combination with other enzymes, e.g., S1 nucleases (Hyrup (1996) *supra*; or as probes or primers for DNA sequence and hybridization (Hyrup (1996) *supra*; Perry-O'Keefe et al. (1996) *Proc. Natl. Acad. Sci. USA* 93: 14670-675).

In another embodiment, PNAs of Tango-77 can be modified, e.g., to enhance their stability or cellular uptake, by attaching lipophilic or other helper groups to PNA, by the formation of PNA-DNA chimeras, or by the use of liposomes or other techniques of drug delivery known in the art. For example, PNA-DNA chimeras of Tango-77 can be generated which may combine the advantageous properties of PNA and DNA. Such chimeras allow DNA recognition enzymes, e.g., RNase H and DNA polymerases, to interact with the DNA portion while the PNA portion

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would provide high binding affinity and specificity. PNA-DNA chimeras can be linked using linkers of appropriate lengths selected in terms of base stacking, number of bonds between the nucleobases, and orientation (Hyrup (1996) *supra*). The synthesis of PNA-DNA chimeras can be performed as described in Hyrup (1996) *supra* and Finn et al. (1996) *Nucleic Acids Res.* 24(17):3357-63. For example, a DNA chain can be synthesized on a solid support using standard phosphoramidite coupling chemistry and modified nucleoside analogs. Compounds such as 5'-(4-methoxytrityl)amino-5'-deoxy-thymidine phosphoramidite can be used as a link between the PNA and the 5' end of DNA (Mag et al. (1989) *Nucleic Acid Res.* 17:5973-88). PNA monomers are then coupled in a stepwise manner to produce a chimeric molecule with a 5' PNA segment and a 3' DNA segment (Finn et al. (1996) *Nucleic Acids Res.* 24(17):3357-63). Alternatively, chimeric molecules can be synthesized with a 5' DNA segment and a 3' PNA segment (Peterser et al. (1975) *Bioorganic Med. Chem. Lett.* 5:1119-11124).

In other embodiments, the oligonucleotide may include other appended groups such as peptides (e.g., for targeting host cell receptors *in vivo*), or agents facilitating transport across the cell membrane (see, e.g., Letsinger et al. (1989) *Proc. Natl. Acad. Sci. USA* 86:6553-6556; Lemaitre et al. (1987) *Proc. Natl. Acad. Sci. USA* 84:648-652; PCT Publication No. WO 88/09810) or the blood-brain barrier (see, e.g., PCT Publication No. WO 89/10134). In addition, oligonucleotides can be modified with hybridization-triggered cleavage agents (see, e.g., Krol et al. (1988) *Bio/Techniques* 6:958-976) or intercalating agents (see, e.g., Zon (1988) *Pharm. Res.* 5:539-549). To this end, the oligonucleotide may be conjugated to another molecule, e.g., a peptide,

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hybridization triggered cross-linking agent, transport agent, hybridization-triggered cleavage agent, etc.

II. Isolated Tango-77 Proteins and Anti-Tango-77 Antibodies

5 One aspect of the invention pertains to isolated Tango-77 proteins, and biologically active portions thereof, as well as polypeptide fragments suitable for use as immunogens to raise anti-Tango-77 antibodies. In one embodiment, native Tango-77 proteins can be isolated
10 from cells or tissue sources by an appropriate purification scheme using standard protein purification techniques. In another embodiment, Tango-77 proteins are produced by recombinant DNA techniques. Alternative to recombinant expression, a Tango-77 protein or polypeptide
15 can be synthesized chemically using standard peptide synthesis techniques.

 An "isolated" or "purified" protein or biologically active portion thereof is substantially free of cellular material or other contaminating proteins from
20 the cell or tissue source from which the Tango-77 protein is derived, or substantially free of chemical precursors or other chemicals when chemically synthesized. The language "substantially free of cellular material" includes preparations of Tango-77 protein in which the
25 protein is separated from cellular components of the cells from which it is isolated or recombinantly produced. Thus, Tango-77 protein that is substantially free of cellular material includes preparations of Tango-77 protein having less than about 30%, 20%, 10%, or
30 5% (by dry weight) of non-Tango-77 protein (also referred to herein as a "contaminating protein"). When the Tango-77 protein or biologically active portion thereof is recombinantly produced, it is also preferably substantially free of culture medium, i.e., culture

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medium represents less than about 20%, 10%, or 5% of the volume of the protein preparation. When Tango-77 protein is produced by chemical synthesis, it is preferably substantially free of chemical precursors or other chemicals, i.e., it is separated from chemical precursors or other chemicals which are involved in the synthesis of the protein. Accordingly such preparations of Tango-77 protein have less than about 30%, 20%, 10%, 5% (by dry weight) of chemical precursors or non-Tango-77 chemicals.

Biologically active portions of a Tango-77 protein include peptides comprising amino acid sequences sufficiently identical to or derived from the amino acid sequence of the Tango-77 protein (e.g., the amino acid sequence shown in SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13), which include fewer amino acids than the full length Tango-77 proteins, and exhibit at least one activity of a Tango-77 protein. Typically, biologically active portions comprise a domain or motif with at least one activity of the Tango-77 protein. A biologically active portion of a Tango-77 protein can be a polypeptide which is, for example, 10, 25, 50, 100 or more amino acids in length.

Moreover, other biologically active portions, in which other regions of the protein are deleted, can be prepared by recombinant techniques and evaluated for one or more of the functional activities of a native Tango-77 protein.

Preferred Tango-77 protein has the amino acid sequence shown of SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13. Other useful Tango-77 proteins are substantially identical to SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13 and retain the functional activity of the protein of SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13 yet differ in

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amino acid sequence due to natural allelic variation or mutagenesis. Accordingly, a useful Tango-77 protein is a protein which includes an amino acid sequence at least about 45%, preferably 55%, 65%, 75%, 85%, 95%, or 99% identical to the amino acid sequence of SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13 and retains the functional activity of the Tango-77 proteins of SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13. In a preferred embodiment, the Tango-77 protein retains a functional activity of the Tango-77 protein of SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11, or SEQ ID NO:13.

To determine the percent identity of two amino acid sequences or of two nucleic acids, the sequences are aligned for optimal comparison purposes (e.g., gaps can be introduced in the sequence of a first amino acid or nucleic acid sequence for optimal alignment with a second amino or nucleic acid sequence). The amino acid residues or nucleotides at corresponding amino acid positions or nucleotide positions are then compared. When a position in the first sequence is occupied by the same amino acid residue or nucleotide as the corresponding position in the second sequence, then the molecules are identical at that position. The percent identity between the two sequences is a function of the number of identical positions shared by the sequences (i.e., % identity = # of identical positions/total # of positions, e.g., overlapping x 100). Preferably, the two sequences are the same length.

The determination of percent homology between two sequences can be accomplished using a mathematical algorithm. A preferred, non-limiting example of a mathematical algorithm utilized for the comparison of two sequences is the algorithm of Karlin and Altschul (1990)

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Proc. Natl. Acad. Sci. USA 87:2264-2268, modified as in Karlin and Altschul (1993) Proc. Natl. Acad. Sci. USA 90:5873-5877. Such an algorithm is incorporated into the NBLAST and XBLAST programs of Altschul, et al. (1990) J. Mol. Biol. 215:403-410. BLAST nucleotide searches can be performed with the NBLAST program, score = 100, wordlength = 12 to obtain nucleotide sequences homologous to Tango-77 nucleic acid molecules of the invention. BLAST protein searches can be performed with the XBLAST program, score = 50, wordlength = 3 to obtain amino acid sequences homologous to Tango-77 protein molecules of the invention. To obtain gapped alignments for comparison purposes, Gapped BLAST can be utilized as described in Altschul et al. (1997) Nucleic Acids Res. 25:3389-3402. When utilizing BLAST and Gapped BLAST programs, the default parameters of the respective programs (e.g., XBLAST and NBLAST) can be used. See <http://www.ncbi.nlm.nih.gov>. Another preferred, non-limiting example of a mathematical algorithm utilized for the comparison of sequences is the algorithm of Myers and Miller, CABIOS (1989). Such an algorithm is incorporated into the ALIGN program (version 2.0) which is part of the GCG sequence alignment software package. When utilizing the ALIGN program for comparing amino acid sequences, a PAM120 weight residue table, a gap length penalty of 12, and a gap penalty of 4 can be used.

The percent identity between two sequences can be determined using techniques similar to those described above, with or without allowing gaps. In calculating percent identity, only exact matches are counted.

The invention also provides Tango-77 chimeric or fusion proteins. As used herein, a Tango-77 "chimeric protein" or "fusion protein" comprises a Tango-77 polypeptide operably linked to a non-Tango-77 polypeptide. A "Tango-77 polypeptide" refers to a

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polypeptide having an amino acid sequence corresponding to Tango-77 polypeptides, whereas a "non-Tango-77 polypeptide" refers to a polypeptide having an amino acid sequence corresponding to a protein which is not substantially identical to the Tango-77 protein, e.g., a protein which is different from the Tango-77 protein and which is derived from the same or a different organism. Within a Tango-77 fusion protein the Tango-77 polypeptide can correspond to all or a portion of a Tango-77 protein, preferably at least one biologically active portion of a Tango-77 protein. Within the fusion protein, the term "operably linked" is intended to indicate that the Tango-77 polypeptide and the non-Tango-77 polypeptide are fused in-frame to each other. The non-Tango-77 polypeptide can be fused to the N-terminus or C-terminus of the Tango-77 polypeptide.

One useful fusion protein is a GST-Tango-77 fusion protein in which the Tango-77 sequences are fused to the C-terminus of the GST sequences. Such fusion proteins can facilitate the purification of recombinant Tango-77.

In another embodiment, the fusion protein is a Tango-77 protein containing a heterologous signal sequence at its N-terminus. For example, the native Tango-77 signal sequence (i.e., about amino acids 1 to 63 of SEQ ID NO:2; SEQ ID NO:4; or about amino acids 1 to 52 of SEQ ID NO:7; SEQ ID NO:8; or about amino acids 1 to 21 of SEQ ID NO:11; SEQ ID NO:12) can be removed and replaced with a signal sequence from another protein. In certain host cells (e.g., mammalian host cells), expression and/or secretion of Tango-77 can be increased through use of a heterologous signal sequence. For example, the gp67 secretory sequence of the baculovirus envelope protein can be used as a heterologous signal sequence (Ausubel et al., supra). Other examples of eukaryotic heterologous signal sequences include the secretory sequences of

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melittin and human placental alkaline phosphatase (Stratagene; La Jolla, California). In yet another example, useful prokaryotic heterologous signal sequences include the phoA secretory signal (Sambrook et al.,
5 supra) and the protein A secretory signal (Pharmacia Biotech; Piscataway, New Jersey).

In yet another embodiment, the fusion protein is an Tango-77-immunoglobulin fusion protein in which all or part of Tango-77 is fused to sequences derived from a
10 member of the immunoglobulin protein family. The Tango-77-immunoglobulin fusion proteins of the invention can be incorporated into pharmaceutical compositions and administered to a subject to inhibit an interaction
15 between a Tango-77 ligand and a Tango-77 receptor on the surface of a cell, to thereby suppress Tango-77-mediated signal transduction in vivo. The Tango-77-immunoglobulin fusion proteins can be used to affect the bioavailability of a Tango-77 cognate ligand. Inhibition of the Tango-77
20 ligand/Tango-77 interaction may be useful therapeutically for both the treatment of inflammatory and autoimmune disorders. Moreover, the Tango-77-immunoglobulin fusion proteins of the invention can be used as immunogens to produce anti-Tango-77 antibodies in a subject, to purify
25 Tango-77 ligands and in screening assays to identify molecules which inhibit the interaction of Tango-77 with a Tango-77 receptor.

Preferably, a Tango-77 chimeric or fusion protein of the invention is produced by standard recombinant DNA techniques. For example, DNA fragments coding for the
30 different polypeptide sequences are ligated together in-frame in accordance with conventional techniques, for example by employing blunt-ended or stagger-ended termini for ligation, restriction enzyme digestion to provide for appropriate termini, filling-in of cohesive ends as
35 appropriate, alkaline phosphatase treatment to avoid

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undesirable joining, and enzymatic ligation. In another embodiment, the fusion gene can be synthesized by conventional techniques including automated DNA synthesizers. Alternatively, PCR amplification of gene fragments can be carried out using anchor primers which give rise to complementary overhangs between two consecutive gene fragments which can subsequently be annealed and reamplified to generate a chimeric gene sequence (see, e.g., *Current Protocols in Molecular Biology*, Ausubel et al. eds., John Wiley & Sons: 1992). Moreover, many expression vectors are commercially available that already encode a fusion moiety (e.g., a GST polypeptide). An Tango-77-encoding nucleic acid can be cloned into such an expression vector such that the fusion moiety is linked in-frame to the Tango-77 protein.

The present invention also pertains to variants of the Tango-77 proteins (i.e., proteins having a sequence which differs from that of the Tango-77 amino acid sequence). Such variants can function as either Tango-77 agonists (mimetics) or as Tango-77 antagonists. Variants of the Tango-77 protein can be generated by mutagenesis, e.g., discrete point mutation or truncation of the Tango-77 protein. An agonist of the Tango-77 protein can retain substantially the same, or a subset, of the biological activities of the naturally occurring form of the Tango-77 protein. An antagonist of the Tango-77 protein can inhibit one or more of the activities of the naturally occurring form of the Tango-77 protein by, for example, competitively binding to a downstream or upstream member of a cellular signaling cascade which includes the Tango-77 protein. Thus, specific biological effects can be elicited by treatment with a variant of limited function. Treatment of a subject with a variant having a subset of the biological activities of the naturally occurring form of the protein can have fewer

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side effects in a subject relative to treatment with the naturally occurring form of the Tango-77 proteins.

Variants of the Tango-77 protein which function as either Tango-77 agonists (mimetics) or as Tango-77
5 antagonists can be identified by screening combinatorial libraries of mutants, e.g., truncation mutants, of the Tango-77 protein for Tango-77 protein agonist or antagonist activity. In one embodiment, a variegated library of Tango-77 variants is generated by
10 combinatorial mutagenesis at the nucleic acid level and is encoded by a variegated gene library. A variegated library of Tango-77 variants can be produced by, for example, enzymatically ligating a mixture of synthetic oligonucleotides into gene sequences such that a
15 degenerate set of potential Tango-77 sequences is expressible as individual polypeptides, or alternatively, as a set of larger fusion proteins (e.g., for phage display) containing the set of Tango-77 sequences therein. There are a variety of methods which can be
20 used to produce libraries of potential Tango-77 variants from a degenerate oligonucleotide sequence. Chemical synthesis of a degenerate gene sequence can be performed in an automatic DNA synthesizer, and the synthetic gene then ligated into an appropriate expression vector. Use
25 of a degenerate set of genes allows for the provision, in one mixture, of all of the sequences encoding the desired set of potential Tango-77 sequences. Methods for synthesizing degenerate oligonucleotides are known in the art (see, e.g., Narang (1983) *Tetrahedron* 39:3; Itakura
30 et al. (1984) *Annu. Rev. Biochem.* 53:323; Itakura et al. (1984) *Science* 198:1056; Ike et al. (1983) *Nucleic Acid Res.* 11:477).

In addition, libraries of fragments of the Tango-77 protein coding sequence can be used to generate
35 a variegated population of Tango-77 fragments for

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screening and subsequent selection of variants of a Tango-77 protein. In one embodiment, a library of coding sequence fragments can be generated by treating a double stranded PCR fragment of a Tango-77 coding sequence with
5 a nuclease under conditions wherein nicking occurs only about once per molecule, denaturing the double stranded DNA, renaturing the DNA to form double stranded DNA which can include sense/antisense pairs from different nicked products, removing single stranded portions from reformed
10 duplexes by treatment with S1 nuclease, and ligating the resulting fragment library into an expression vector. By this method, an expression library can be derived which encodes N-terminal and internal fragments of various sizes of the Tango-77 protein.

15 Several techniques are known in the art for screening gene products of combinatorial libraries made by point mutations or truncation, and for screening cDNA libraries for gene products having a selected property. Such techniques are adaptable for rapid screening of the
20 gene libraries generated by the combinatorial mutagenesis of Tango-77 proteins. The most widely used techniques, which are amenable to high through-put analysis, for screening large gene libraries typically include cloning the gene library into replicable expression vectors,
25 transforming appropriate cells with the resulting library of vectors, and expressing the combinatorial genes under conditions in which detection of a desired activity facilitates isolation of the vector encoding the gene whose product was detected. Recursive ensemble
30 mutagenesis (REM), a technique which enhances the frequency of functional mutants in the libraries, can be used in combination with the screening assays to identify Tango-77 variants (Arkin and Yourvan (1992) *Proc. Natl. Acad. Sci. USA* 89:7811-7815; Delgrave et al. (1993)
35 *Protein Engineering* 6(3):327-331).

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An isolated Tango-77 protein, or a portion or fragment thereof, can be used as an immunogen to generate antibodies that bind Tango-77 using standard techniques for polyclonal and monoclonal antibody preparation. The
5 full-length Tango-77 protein can be used or, alternatively, the invention provides antigenic peptide fragments of Tango-77 for use as immunogens. The antigenic peptide of Tango-77 comprises at least 8 (preferably 10, 15, 20, or 30) amino acid residues of the
10 amino acid sequence shown in SEQ ID NO:2, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:9, SEQ ID NO:11 or SEQ ID NO:13 and encompasses an epitope of Tango-77 such that an antibody raised against the peptide forms a specific immune complex with Tango-77.

15 A Tango-77 immunogen typically is used to prepare antibodies by immunizing a suitable subject (e.g., rabbit, goat, mouse or other mammal) with the immunogen. An appropriate immunogenic preparation can contain, for example, recombinantly expressed Tango-77 protein or a
20 chemically synthesized Tango-77 polypeptide. The preparation can further include an adjuvant, such as Freund's complete or incomplete adjuvant, or similar immunostimulatory agent. Immunization of a suitable subject with an immunogenic Tango-77 preparation induces
25 a polyclonal anti-Tango-77 antibody response.

Accordingly, another aspect of the invention pertains to anti-Tango-77 antibodies. The term "antibody" as used herein refers to immunoglobulin molecules and immunologically active portions of
30 immunoglobulin molecules, i.e., molecules that contain an antigen binding site which specifically binds an antigen, such as Tango-77. A molecule which specifically binds to Tango-77 is a molecule which binds Tango-77, but does not substantially bind other molecules in a sample, e.g., a
35 biological sample, which naturally contains Tango-77.

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Examples of immunologically active portions of immunoglobulin molecules include F(ab) and F(ab')₂ fragments which can be generated by treating the antibody with an enzyme such as pepsin. The invention provides polyclonal and monoclonal antibodies that bind Tango-77. The term "monoclonal antibody" or "monoclonal antibody composition", as used herein, refers to a population of antibody molecules that contain only one species of an antigen binding site capable of immunoreacting with a particular epitope of Tango-77. A monoclonal antibody composition thus typically displays a single binding affinity for a particular Tango-77 protein with which it immunoreacts.

Polyclonal anti-Tango-77 antibodies can be prepared as described above by immunizing a suitable subject with a Tango-77 immunogen. The anti-Tango-77 antibody titer in the immunized subject can be monitored over time by standard techniques, such as with an enzyme linked immunosorbent assay (ELISA) using immobilized Tango-77. If desired, the antibody molecules directed against Tango-77 can be isolated from the mammal (e.g., from the blood) and further purified by well-known techniques, such as protein A chromatography to obtain the IgG fraction. At an appropriate time after immunization, e.g., when the anti-Tango-77 antibody titers are highest, antibody-producing cells can be obtained from the subject and used to prepare monoclonal antibodies by standard techniques, such as the hybridoma technique originally described by Kohler and Milstein (1975) *Nature* 256:495-497, the human B cell hybridoma technique (Kozbor et al. (1983) *Immunol Today* 4:72), the EBV-hybridoma technique (Cole et al. (1985), *Monoclonal Antibodies and Cancer Therapy*, Alan R. Liss, Inc., pp. 77-96) or trioma techniques. The technology for producing hybridomas is well known (see generally Current

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Protocols in Immunology (1994) Coligan et al. (eds.) John Wiley & Sons, Inc., New York, NY). Briefly, an immortal cell line (typically a myeloma) is fused to lymphocytes (typically splenocytes) from a mammal immunized with a
5 Tango-77 immunogen as described above, and the culture supernatants of the resulting hybridoma cells are screened to identify a hybridoma producing a monoclonal antibody that binds Tango-77.

Any of the many well known protocols used for
10 fusing lymphocytes and immortalized cell lines can be applied for the purpose of generating an anti-Tango-77 monoclonal antibody (see, e.g., Current Protocols in Immunology, supra; Galfre et al. (1977) Nature 266:55052; R.H. Kenneth, in Monoclonal Antibodies: A New Dimension
15 In Biological Analyses, Plenum Publishing Corp., New York, New York (1980); and Lerner (1981) Yale J. Biol. Med., 54:387-402. Moreover, the ordinarily skilled worker will appreciate that there are many variations of such methods which also would be useful. Typically, the
20 immortal cell line (e.g., a myeloma cell line) is derived from the same mammalian species as the lymphocytes. For example, murine hybridomas can be made by fusing lymphocytes from a mouse immunized with an immunogenic preparation of the present invention with an immortalized
25 mouse cell line, e.g., a myeloma cell line that is sensitive to culture medium containing hypoxanthine, aminopterin and thymidine ("HAT medium"). Any of a number of myeloma cell lines can be used as a fusion partner according to standard techniques, e.g., the P3-
30 NS1/1-Ag4-1, P3-x63-Ag8.653 or Sp2/O-Ag14 myeloma lines. These myeloma lines are available from ATCC. Typically, HAT-sensitive mouse myeloma cells are fused to mouse splenocytes using polyethylene glycol ("PEG"). Hybridoma cells resulting from the fusion are then selected using
35 HAT medium, which kills unfused and unproductively fused

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myeloma cells (unfused splenocytes die after several days because they are not transformed). Hybridoma cells producing a monoclonal antibody of the invention are detected by screening the hybridoma culture supernatants
5 for antibodies that bind Tango-77, e.g., using a standard ELISA assay.

Alternative to preparing monoclonal antibody-secreting hybridomas, a monoclonal anti-Tango-77 antibody can be identified and isolated by screening a recombinant
10 combinatorial immunoglobulin library (e.g., an antibody phage display library) with Tango-77 to thereby isolate immunoglobulin library members that bind Tango-77. Kits for generating and screening phage display libraries are commercially available (e.g., the Pharmacia Recombinant
15 Phage Antibody System, Catalog No. 27-9400-01; and the Stratagene SurfZAP™ Phage Display Kit, Catalog No. 240612). Additionally, examples of methods and reagents particularly amenable for use in generating and screening antibody display library can be found in, for example,
20 U.S. Patent No. 5,223,409; PCT Publication No. WO 92/18619; PCT Publication No. WO 91/17271; PCT Publication No. WO 92/20791; PCT Publication No. WO 92/15679; PCT Publication No. WO 93/01288; PCT Publication No. WO 92/01047; PCT Publication No. WO
25 92/09690; PCT Publication No. WO 90/02809; Fuchs et al. (1991) *Bio/Technology* 9:1370-1372; Hay et al. (1992) *Hum. Antibod. Hybridomas* 3:81-85; Huse et al. (1989) *Science* 246:1275-1281; Griffiths et al. (1993) *EMBO J* 12:725-734.

Additionally, recombinant anti-Tango-77
30 antibodies, such as chimeric and humanized monoclonal antibodies, comprising both human and non-human portions, which can be made using standard recombinant DNA techniques, are within the scope of the invention. Such chimeric and humanized monoclonal antibodies can be
35 produced by recombinant DNA techniques known in the art,

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- for example using methods described in PCT Publication No. WO 87/02671; European Patent Application 184,187; European Patent Application 171,496; European Patent Application 173,494; PCT Publication No. WO 86/01533;
- 5 U.S. Patent No. 4,816,567; European Patent Application 125,023; Better et al. (1988) *Science* 240:1041-1043; Liu et al. (1987) *Proc. Natl. Acad. Sci. USA* 84:3439-3443; Liu et al. (1987) *J. Immunol.* 139:3521-3526; Sun et al. (1987) *Proc. Natl. Acad. Sci. USA* 84:214-218; Nishimura
- 10 et al. (1987) *Canc. Res.* 47:999-1005; Wood et al. (1985) *Nature* 314:446-449; and Shaw et al. (1988) *J. Natl. Cancer Inst.* 80:1553-1559; Morrison (1985) *Science* 229:1202-1207; Oi et al. (1986) *Bio/Techniques* 4:214; U.S. Patent 5,225,539; Jones et al. (1986) *Nature*
- 15 321:552-525; Verhoeyan et al. (1988) *Science* 239:1534; and Beidler et al. (1988) *J. Immunol.* 141:4053-4060.

Completely human antibodies are particularly desirable for therapeutic treatment of human patients. Such antibodies can be produced using transgenic mice

20 which are incapable of expressing endogenous immunoglobulin heavy and light chains genes, but which can express human heavy and light chain genes. The transgenic mice are immunized in the normal fashion with a selected antigen, e.g., all or a portion of Tango-77.

25 Monoclonal antibodies directed against the antigen can be obtained using conventional hybridoma technology. The human immunoglobulin transgenes harbored by the transgenic mice rearrange during B cell differentiation, and subsequently undergo class switching and somatic

30 mutation. Thus, using such a technique, it is possible to produce therapeutically useful IgG, IgA and IgE antibodies. For an overview of this technology for producing human antibodies, see Lonberg and Huszar (1995, *Int. Rev. Immunol.* 13:65-93). For a detailed discussion

35 of this technology for producing human antibodies and

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human monoclonal antibodies and protocols for producing such antibodies, see, e.g., U.S. Patent 5,625,126; U.S. Patent 5,633,425; U.S. Patent 5,569,825; U.S. Patent 5,661,016; and U.S. Patent 5,545,806. In addition, 5 companies such as Abgenix, Inc. (Freemont, CA), can be engaged to provide human antibodies directed against a selected antigen using technology similar to the described above.

Completely human antibodies which recognize a 10 selected epitope can be generated using a technique referred to as "guided selection." In this approach a selected non-human monoclonal antibody, e.g., a murine antibody, is used to guide the selection of a completely human antibody recognizing the same epitope.

15 First, a non-human monoclonal antibody which binds a selected antigen (epitope), e.g., an antibody which inhibits Tango-77 activity, is identified. The heavy chain and the light chain of the non-human antibody are cloned and used to create phage display Fab fragments. 20 For example, the heavy chain gene can be cloned into a plasmid vector so that the heavy chain can be secreted from bacteria. The light chain gene can be cloned into a phage coat protein gene so that the light chain can be expressed on the surface of phage. A repertoire (random 25 collection) of human light chains fused to phage is used to infect the bacteria which express the non-human heavy chain. The resulting progeny phage display hybrid antibodies (human light chain/non-human heavy chain). The selected antigen is used in a panning screen to 30 select phage which bind the selected antigen. Several rounds of selection may be required to identify such phage. Next, human light chain genes are isolated from the selected phage which bind the selected antigen. These selected human light chain genes are then used to 35 guide the selection of human heavy chain genes as

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follows. The selected human light chain genes are inserted into vectors for expression by bacteria. Bacteria expressing the selected human light chains are infected with a repertoire of human heavy chains fused to phage. The resulting progeny phage display human antibodies (human light chain/human heavy chain).

Next, the selected antigen is used in a panning screen to select phage which bind the selected antigen. The phage selected in this step display completely human antibody which recognize the same epitope recognized by the original selected, non-human monoclonal antibody. The genes encoding both the heavy and light chains are readily isolated and be further manipulated for production of human antibody. This technology is described by Jespers et al. (1994, *Bio/technology* 12:899-903).

An anti-Tango-77 antibody (e.g., monoclonal antibody) can be used to isolate Tango-77 by standard techniques, such as affinity chromatography or immunoprecipitation. An anti-Tango-77 antibody can facilitate the purification of natural Tango-77 from cells and of recombinantly produced Tango-77 expressed in host cells. Moreover, an anti-Tango-77 antibody can be used to detect Tango-77 protein (e.g., in a cellular lysate or cell supernatant) in order to evaluate the abundance and pattern of expression of the Tango-77 protein. Anti-Tango-77 antibodies can be used diagnostically to monitor protein levels in tissue as part of a clinical testing procedure, e.g., to, for example, determine the efficacy of a given treatment regimen. Detection can be facilitated by coupling the antibody to a detectable substance. Examples of detectable substances include various enzymes, prosthetic groups, fluorescent materials, luminescent materials, bioluminescent materials, and radioactive materials.

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Examples of suitable enzymes include horseradish peroxidase, alkaline phosphatase, β -galactosidase, or acetylcholinesterase; examples of suitable prosthetic group complexes include streptavidin/biotin and
5 avidin/biotin; examples of suitable fluorescent materials include umbelliferone, fluorescein, fluorescein isothiocyanate, rhodamine, dichlorotriazinylamine fluorescein, dansyl chloride or phycoerythrin; an example of a luminescent material includes luminol; examples of
10 bioluminescent materials include luciferase, luciferin, and aequorin, and examples of suitable radioactive material include ^{125}I , ^{131}I , ^{35}S or ^3H .

III. Recombinant Expression Vectors and Host Cells

Another aspect of the invention pertains to
15 vectors, preferably expression vectors, containing a nucleic acid molecule encoding Tango-77 (or a portion thereof). As used herein, the term "vector" refers to a nucleic acid molecule capable of transporting another nucleic acid to which it has been linked. One type of
20 vector is a "plasmid", which refers to a circular double stranded DNA loop into which additional DNA segments can be ligated. Another type of vector is a viral vector, wherein additional DNA segments can be ligated into the viral genome. Certain vectors are capable of autonomous
25 replication in a host cell into which they are introduced (e.g., bacterial vectors having a bacterial origin of replication and episomal mammalian vectors). Other vectors (e.g., non-episomal mammalian vectors) are integrated into the genome of a host cell upon
30 introduction into the host cell, and thereby are replicated along with the host genome. Moreover, certain vectors, expression vectors, are capable of directing the expression of genes to which they are operably linked. In general, expression vectors of utility in recombinant

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DNA techniques are often in the form of plasmids (vectors). However, the invention is intended to include such other forms of expression vectors, such as viral vectors (e.g., replication defective retroviruses, adenoviruses and adeno-associated viruses), which serve equivalent functions.

The recombinant expression vectors of the invention comprise a nucleic acid of the invention in a form suitable for expression of the nucleic acid in a host cell, which means that the recombinant expression vectors include one or more regulatory sequences, selected on the basis of the host cells to be used for expression, which is operably linked to the nucleic acid sequence to be expressed. Within a recombinant expression vector, "operably linked" is intended to mean that the nucleotide sequence of interest is linked to the regulatory sequence(s) in a manner which allows for expression of the nucleotide sequence (e.g., in an *in vitro* transcription/translation system or in a host cell when the vector is introduced into the host cell). The term "regulatory sequence" is intended to include promoters, enhancers and other expression control elements (e.g., polyadenylation signals). Such regulatory sequences are described, for example, in Goeddel; *Gene Expression Technology: Methods in Enzymology* 185, Academic Press, San Diego, CA (1990). Regulatory sequences include those which direct constitutive expression of a nucleotide sequence in many types of host cell and those which direct expression of the nucleotide sequence only in certain host cells (e.g., tissue-specific regulatory sequences). It will be appreciated by those skilled in the art that the design of the expression vector can depend on such factors as the choice of the host cell to be transformed, the level of expression of protein desired, etc. The expression

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vectors of the invention can be introduced into host cells to thereby produce proteins or peptides, including fusion proteins or peptides, encoded by nucleic acids as described herein (e.g., Tango-77 proteins, mutant forms of Tango-77, fusion proteins, etc.).

The recombinant expression vectors of the invention can be designed for expression of Tango-77 in prokaryotic or eukaryotic cells, e.g., bacterial cells such as *E. coli*, insect cells (using baculovirus expression vectors), yeast cells or mammalian cells. Suitable host cells are discussed further in Goeddel, *Gene Expression Technology: Methods in Enzymology* 185, Academic Press, San Diego, CA (1990). Alternatively, the recombinant expression vector can be transcribed and translated *in vitro*, for example using T7 promoter regulatory sequences and T7 polymerase.

Expression of proteins in prokaryotes is most often carried out in *E. coli* with vectors containing constitutive or inducible promoters directing the expression of either fusion or non-fusion proteins. Fusion vectors add a number of amino acids to a protein encoded therein, usually to the amino terminus of the recombinant protein. Such fusion vectors typically serve three purposes: 1) to increase expression of recombinant protein; 2) to increase the solubility of the recombinant protein; and 3) to aid in the purification of the recombinant protein by acting as a ligand in affinity purification. Often, in fusion expression vectors, a proteolytic cleavage site is introduced at the junction of the fusion moiety and the recombinant protein to enable separation of the recombinant protein from the fusion moiety subsequent to purification of the fusion protein. Such enzymes, and their cognate recognition sequences, include Factor Xa, thrombin and enterokinase. Typical fusion expression vectors include pGEX (Pharmacia

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Biotech Inc; Smith and Johnson (1988) *Gene* 67:31-40), pMAL (New England Biolabs, Beverly, MA) and pRIT5 (Pharmacia, Piscataway, NJ) which fuse glutathione S-transferase (GST), maltose E binding protein, or protein 5 A, respectively, to the target recombinant protein.

Examples of suitable inducible non-fusion *E. coli* expression vectors include pTrc (Amann et al. (1988) *Gene* 69:301-315) and pET 11d (Studier et al., *Gene Expression Technology: Methods in Enzymology* 185, Academic Press, 10 San Diego, California (1990) 60-89). Target gene expression from the pTrc vector relies on host RNA polymerase transcription from a hybrid trp-lac fusion promoter. Target gene expression from the pET 11d vector relies on transcription from a T7 *gn10*-lac fusion 15 promoter mediated by a coexpressed viral RNA polymerase (T7 *gn1*). This viral polymerase is supplied by host strains BL21(DE3) or HMS174(DE3) from a resident λ prophage harboring a T7 *gn1* gene under the transcriptional control of the *lacUV 5* promoter.

20 One strategy to maximize recombinant protein expression in *E. coli* is to express the protein in a host bacteria with an impaired capacity to proteolytically cleave the recombinant protein (Gottesman, *Gene Expression Technology: Methods in Enzymology* 185, 25 Academic Press, San Diego, California (1990) 119-128). Another strategy is to alter the nucleic acid sequence of the nucleic acid to be inserted into an expression vector so that the individual codons for each amino acid are those preferentially utilized in *E. coli* (Wada et al. 30 (1992) *Nucleic Acids Res.* 20:2111-2118). Such alteration of nucleic acid sequences of the invention can be carried out by standard DNA synthesis techniques.

In another embodiment, the Tango-77 expression vector is a yeast expression vector. Examples of vectors 35 for expression in yeast *S. cerevisiae* include pYepSec1

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(Baldari et al. (1987) *EMBO J.* 6:229-234), pMFa (Kurjan and Herskowitz (1982) *Cell* 30:933-943), pJRY88 (Schultz et al. (1987) *Gene* 54:113-123), pYES2 (Invitrogen Corporation, San Diego, CA), and picZ (Invitrogen Corp, 5 San Diego, CA).

Alternatively, Tango-77 can be expressed in insect cells using baculovirus expression vectors. Baculovirus vectors available for expression of proteins in cultured insect cells (e.g., Sf 9 cells) include the pAc series 10 (Smith et al. (1983) *Mol. Cell Biol.* 3:2156-2165) and the pVL series (Lucklow and Summers (1989) *Virology* 170:31-39).

In yet another embodiment, a nucleic acid of the invention is expressed in mammalian cells using a 15 mammalian expression vector. Examples of mammalian expression vectors include pCDM8 (Seed (1987) *Nature* 329:840) and pMT2PC (Kaufman et al. (1987) *EMBO J.* 6:187-195). When used in mammalian cells, the expression vector's control functions are often provided by viral 20 regulatory elements. For example, commonly used promoters are derived from polyoma, Adenovirus 2, cytomegalovirus and Simian Virus 40. For other suitable expression systems for both prokaryotic and eukaryotic cells see chapters 16 and 17 of Sambrook et al. (*supra*).

25 In another embodiment, the recombinant mammalian expression vector is capable of directing expression of the nucleic acid preferentially in a particular cell type (e.g., tissue-specific regulatory elements are used to express the nucleic acid). Tissue-specific regulatory 30 elements are known in the art. Non-limiting examples of suitable tissue-specific promoters include the albumin promoter (liver-specific; Pinkert et al. (1987) *Genes Dev.* 1:268-277), lymphoid-specific promoters (Calame and Eaton (1988) *Adv. Immunol.* 43:235-275), in particular 35 promoters of T cell receptors (Winoto and Baltimore

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(1989) *EMBO J.* 8:729-733) and immunoglobulins (Banerji et al. (1983) *Cell* 33:729-740; Queen and Baltimore (1983) *Cell* 33:741-748), neuron-specific promoters (e.g., the neurofilament promoter; Byrne and Ruddle (1989) *Proc. Natl. Acad. Sci. USA* 86:5473-5477), pancreas-specific promoters (Edlund et al. (1985) *Science* 230:912-916), and mammary gland-specific promoters (e.g., milk whey promoter; U.S. Patent No. 4,873,316 and European Application Publication No. 264,166). Developmentally-regulated promoters are also encompassed, for example the murine hox promoters (Kessel and Gruss (1990) *Science* 249:374-379) and the α -fetoprotein promoter (Campes and Tilghman (1989) *Genes Dev.* 3:537-546).

The invention further provides a recombinant expression vector comprising a DNA molecule of the invention cloned into the expression vector in an antisense orientation. That is, the DNA molecule is operably linked to a regulatory sequence in a manner which allows for expression (by transcription of the DNA molecule) of an RNA molecule which is antisense to Tango-77 mRNA. Regulatory sequences operably linked to a nucleic acid cloned in the antisense orientation can be chosen which direct the continuous expression of the antisense RNA molecule in a variety of cell types, for instance viral promoters and/or enhancers, or regulatory sequences can be chosen which direct constitutive, tissue specific or cell type specific expression of antisense RNA. The antisense expression vector can be in the form of a recombinant plasmid, phagemid or attenuated virus in which antisense nucleic acids are produced under the control of a high efficiency regulatory region, the activity of which can be determined by the cell type into which the vector is introduced. For a discussion of the regulation of gene expression using antisense genes see

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Weintraub et al. (*Reviews - Trends in Genetics*, Vol. 1(1) 1986).

Another aspect of the invention pertains to host cells into which a recombinant expression vector of the invention has been introduced. The terms "host cell" and "recombinant host cell" are used interchangeably herein. It is understood that such terms refer not only to the particular subject cell but to the progeny or potential progeny of such a cell. Because certain modifications may occur in succeeding generations due to either mutation or environmental influences, such progeny may not, in fact, be identical to the parent cell, but are still included within the scope of the term as used herein.

A host cell can be any prokaryotic or eukaryotic cell. For example, Tango-77 protein can be expressed in bacterial cells such as *E. coli*, insect cells, yeast or mammalian cells (such as Chinese hamster ovary cells (CHO) or COS cells). Other suitable host cells are known to those skilled in the art.

Vector DNA can be introduced into prokaryotic or eukaryotic cells via conventional transformation or transfection techniques. As used herein, the terms "transformation" and "transfection" are intended to refer to a variety of art-recognized techniques for introducing foreign nucleic acid (e.g., DNA) into a host cell, including calcium phosphate or calcium chloride coprecipitation, DEAE-dextran-mediated transfection, lipofection, or electroporation. Suitable methods for transforming or transfecting host cells can be found in Sambrook, et al. (*supra*), and other laboratory manuals.

For stable transfection of mammalian cells, it is known that, depending upon the expression vector and transfection technique used, only a small fraction of cells may integrate the foreign DNA into their genome.

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In order to identify and select these integrants, a gene that encodes a selectable marker (e.g., for resistance to antibiotics) is generally introduced into the host cells along with the gene of interest. Preferred selectable
5 markers include those which confer resistance to drugs, such as G418, hygromycin and methotrexate. Nucleic acid encoding a selectable marker can be introduced into a host cell on the same vector as that encoding Tango-77 or can be introduced on a separate vector. Cells stably
10 transfected with the introduced nucleic acid can be identified by drug selection (e.g., cells that have incorporated the selectable marker gene will survive, while the other cells die).

A host cell of the invention, such as a
15 prokaryotic or eukaryotic host cell in culture, can be used to produce (i.e., express) Tango-77 protein. Accordingly, the invention further provides methods for producing Tango-77 protein using the host cells of the invention. In one embodiment, the method comprises
20 culturing the host cell of invention (into which a recombinant expression vector encoding Tango-77 has been introduced) in a suitable medium such that Tango-77 protein is produced. In another embodiment, the method further comprises isolating Tango-77 from the medium or
25 the host cell.

The host cells of the invention can also be used to produce nonhuman transgenic animals. For example, in one embodiment, a host cell of the invention is a fertilized oocyte or an embryonic stem cell into which
30 Tango-77-coding sequences have been introduced. Such host cells can then be used to create non-human transgenic animals in which exogenous Tango-77 sequences have been introduced into their genome or homologous recombinant animals in which endogenous Tango-77
35 sequences have been altered. Such animals are useful for

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studying the function and/or activity of Tango-77 and for identifying and/or evaluating modulators of Tango-77 activity. As used herein, a "transgenic animal" is a non-human animal, preferably a mammal, more preferably a rodent such as a rat or mouse, in which one or more of the cells of the animal includes a transgene. Other examples of transgenic animals include non-human primates, sheep, dogs, cows, goats, chickens, amphibians, etc. A transgene is exogenous DNA which is integrated into the genome of a cell from which a transgenic animal develops and which remains in the genome of the mature animal, thereby directing the expression of an encoded gene product in one or more cell types or tissues of the transgenic animal. As used herein, an "homologous recombinant animal" is a non-human animal, preferably a mammal, more preferably a mouse, in which an endogenous Tango-77 gene has been altered by homologous recombination between the endogenous gene and an exogenous DNA molecule introduced into a cell of the animal, e.g., an embryonic cell of the animal, prior to development of the animal.

A transgenic animal of the invention can be created by introducing Tango-77-encoding nucleic acid into the male pronuclei of a fertilized oocyte, e.g., by microinjection, retroviral infection, and allowing the oocyte to develop in a pseudopregnant female foster animal. The Tango-77 cDNA sequence e.g., that of (SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6; SEQ ID NO:10 or the cDNA of ATCC 98807) can be introduced as a transgene into the genome of a non-human animal. Alternatively, a nonhuman homologue of the human Tango-77 gene, such as a mouse Tango-77 gene, can be isolated based on hybridization to the human Tango-77 cDNA and used as a transgene. Intronic sequences and polyadenylation signals can also be included in the transgene to increase the efficiency

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of expression of the transgene. A tissue-specific regulatory sequence(s) can be operably linked to the Tango-77 transgene to direct expression of Tango-77 protein to particular cells. Methods for generating transgenic animals via embryo manipulation and microinjection, particularly animals such as mice, have become conventional in the art and are described, for example, in U.S. Patent Nos. 4,736,866 and 4,870,009, U.S. Patent No. 4,873,191 and in Hogan, *Manipulating the Mouse Embryo* (Cold Spring Harbor Laboratory Press, Cold Spring Harbor, N.Y., 1986). Similar methods are used for production of other transgenic animals. A transgenic founder animal can be identified based upon the presence of the Tango-77 transgene in its genome and/or expression of Tango-77 mRNA in tissues or cells of the animals. A transgenic founder animal can then be used to breed additional animals carrying the transgene. Moreover, transgenic animals carrying a transgene encoding Tango-77 can further be bred to other transgenic animals carrying other transgenes.

To create an homologous recombinant animal, a vector is prepared which contains at least a portion of a Tango-77 gene (e.g., a human or a non-human homolog of the Tango-77 gene, e.g., a murine Tango-77 gene) into which a deletion, addition or substitution has been introduced to thereby alter, e.g., functionally disrupt, the Tango-77 gene. In a preferred embodiment, the vector is designed such that, upon homologous recombination, the endogenous Tango-77 gene is functionally disrupted (i.e., no longer encodes a functional protein; also referred to as a "knock out" vector). Alternatively, the vector can be designed such that, upon homologous recombination, the endogenous Tango-77 gene is mutated or otherwise altered but still encodes functional protein (e.g., the upstream regulatory region can be altered to thereby

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alter the expression of the endogenous Tango-77 protein). In the homologous recombination vector, the altered portion of the Tango-77 gene is flanked at its 5' and 3' ends by additional nucleic acid of the Tango-77 gene to
5 allow for homologous recombination to occur between the exogenous Tango-77 gene carried by the vector and an endogenous Tango-77 gene in an embryonic stem cell. The additional flanking Tango-77 nucleic acid is of sufficient length for successful homologous recombination
10 with the endogenous gene. Typically, several kilobases of flanking DNA (both at the 5' and 3' ends) are included in the vector (see, e.g., Thomas and Capecchi (1987) *Cell* 51:503 for a description of homologous recombination vectors). The vector is introduced into an embryonic
15 stem cell line (e.g., by electroporation) and cells in which the introduced Tango-77 gene has homologously recombined with the endogenous Tango-77 gene are selected (see, e.g., Li et al. (1992) *Cell* 69:915). The selected cells are then injected into a blastocyst of an animal
20 (e.g., a mouse) to form aggregation chimeras (see, e.g., Bradley in *Teratocarcinomas and Embryonic Stem Cells: A Practical Approach*, Robertson, ed. (IRL, Oxford, 1987) pp. 113-152). A chimeric embryo can then be implanted into a suitable pseudopregnant female foster animal and
25 the embryo brought to term. Progeny harboring the homologously recombined DNA in their germ cells can be used to breed animals in which all cells of the animal contain the homologously recombined DNA by germline transmission of the transgene. Methods for constructing
30 homologous recombination vectors and homologous recombinant animals are described further in Bradley (1991) *Current Opinion in Bio/Technology* 2:823-829 and in PCT Publication Nos. WO 90/11354, WO 91/01140, WO 92/0968, and WO 93/04169.

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In another embodiment, transgenic non-human animals can be produced which contain selected systems which allow for regulated expression of the transgene. One example of such a system is the cre/loxP recombinase system of bacteriophage P1. For a description of the cre/loxP recombinase system, see, e.g., Lakso et al. (1992) *Proc. Natl. Acad. Sci. USA* 89:6232-6236. Another example of a recombinase system is the FLP recombinase system of *Saccharomyces cerevisiae* (O'Gorman et al. (1991) *Science* 251:1351-1355. If a cre/loxP recombinase system is used to regulate expression of the transgene, animals containing transgenes encoding both the Cre recombinase and a selected protein are required. Such animals can be provided through the construction of "double" transgenic animals, e.g., by mating two transgenic animals, one containing a transgene encoding a selected protein and the other containing a transgene encoding a recombinase.

Clones of the non-human transgenic animals described herein can also be produced according to the methods described in Wilmut et al. (1997) *Nature* 385:810-813 and PCT Publication Nos. WO 97/07668 and WO 97/07669. In brief, a cell, e.g., a somatic cell, from the transgenic animal can be isolated and induced to exit the growth cycle and enter G₀ phase. The quiescent cell can then be fused, e.g., through the use of electrical pulses, to an enucleated oocyte from an animal of the same species from which the quiescent cell is isolated. The reconstructed oocyte is then cultured such that it develops to morula or blastocyte and then transferred to pseudopregnant female foster animal. The offspring borne of this female foster animal will be a clone of the animal from which the cell, e.g., the somatic cell, is isolated.

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IV. Pharmaceutical Compositions

The Tango-77 nucleic acid molecules, Tango-77 proteins, and anti-Tango-77 antibodies (also referred to herein as "active compounds") of the invention can be
5 incorporated into pharmaceutical compositions suitable for administration. Such compositions typically comprise the nucleic acid molecule, protein, or antibody and a pharmaceutically acceptable carrier. As used herein the language "pharmaceutically acceptable carrier" is
10 intended to include any and all solvents, dispersion media, coatings, antibacterial and antifungal agents, isotonic and absorption delaying agents, and the like, compatible with pharmaceutical administration. The use of such media and agents for pharmaceutically active
15 substances is well known in the art. Except insofar as any conventional media or agent is incompatible with the active compound, use thereof in the compositions is contemplated. Supplementary active compounds can also be incorporated into the compositions.

20 A pharmaceutical composition of the invention is formulated to be compatible with its intended route of administration. Examples of routes of administration include parenteral, (e.g. intravenous, intradermal, subcutaneous) (e.g., oral inhalation), transdermal
25 (topical), transmucosal, and rectal administration. Solutions or suspensions used for parenteral, intradermal, or subcutaneous application can include the following components: a sterile diluent such as water for injection, saline solution, fixed oils, polyethylene
30 glycols, glycerine, propylene glycol or other synthetic solvents; antibacterial agents such as benzyl alcohol or methyl parabens; antioxidants such as ascorbic acid or sodium bisulfite; chelating agents such as ethylenediaminetetraacetic acid; buffers such as
35 acetates, citrates or phosphates and agents for the

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adjustment of tonicity such as sodium chloride or dextrose. pH can be adjusted with acids or bases, such as hydrochloric acid or sodium hydroxide. The parenteral preparation can be enclosed in ampoules, disposable
5 syringes or multiple dose vials made of glass or plastic.

Pharmaceutical compositions suitable for injectable use include sterile aqueous solutions (where water soluble) or dispersions and sterile powders for the extemporaneous preparation of sterile injectable
10 solutions or dispersions. For intravenous administration, suitable carriers include physiological saline, bacteriostatic water, Cremophor EL™ (BASF; Parsippany, NJ) or phosphate buffered saline (PBS). In all cases, the composition must be sterile and should be
15 fluid to the extent that easy syringability exists. It must be stable under the conditions of manufacture and storage and must be preserved against the contaminating action of microorganisms such as bacteria and fungi. The carrier can be a solvent or dispersion medium containing,
20 for example, water, ethanol, polyol (for example, glycerol, propylene glycol, and liquid polyethylene glycol, and the like), and suitable mixtures thereof. The proper fluidity can be maintained, for example, by the use of a coating such as lecithin, by the maintenance
25 of the required particle size in the case of dispersion and by the use of surfactants. Prevention of the action of microorganisms can be achieved by various antibacterial and antifungal agents, for example, parabens, chlorobutanol, phenol, ascorbic acid,
30 thimerosal, and the like. In many cases, it will be preferable to include isotonic agents, for example, sugars, polyalcohols such as mannitol, sorbitol, sodium chloride in the composition. Prolonged absorption of the injectable compositions can be brought about by including

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in the composition an agent which delays absorption, for example, aluminum monostearate and gelatin.

Sterile injectable solutions can be prepared by incorporating the active compound (e.g., a Tango-77 protein or anti-Tango-77 antibody) in the required amount in an appropriate solvent with one or a combination of ingredients enumerated above, as required, followed by filtered sterilization. Generally, dispersions are prepared by incorporating the active compound into a sterile vehicle which contains a basic dispersion medium and the required other ingredients from those enumerated above. In the case of sterile powders for the preparation of sterile injectable solutions, the preferred methods of preparation are vacuum drying and freeze-drying which yields a powder of the active ingredient plus any additional desired ingredient from a previously sterile-filtered solution thereof.

Oral compositions generally include an inert diluent or an edible carrier. They can be enclosed in gelatin capsules or compressed into tablets. For the purpose of oral therapeutic administration, the active compound can be incorporated with excipients and used in the form of tablets, troches, or capsules. Oral compositions can also be prepared using a fluid carrier for use as a mouthwash, wherein the compound in the fluid carrier is applied orally and swished and expectorated or swallowed. Pharmaceutically compatible binding agents, and/or adjuvant materials can be included as part of the composition. The tablets, pills, capsules, troches and the like can contain any of the following ingredients, or compounds of a similar nature: a binder such as microcrystalline cellulose, gum tragacanth or gelatin; an excipient such as starch or lactose, a disintegrating agent such as alginic acid, Primogel, or corn starch; a lubricant such as magnesium stearate or Sterotes; a

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glidant such as colloidal silicon dioxide; a sweetening agent such as sucrose or saccharin; or a flavoring agent such as peppermint, methyl salicylate, or orange flavoring.

5 For administration by inhalation, the compounds are delivered in the form of an aerosol spray from a pressurized container or dispenser which contains a suitable propellant, e.g., a gas such as carbon dioxide, or a nebulizer.

10 Systemic administration can also be by transmucosal or transdermal means. For transmucosal or transdermal administration, penetrants appropriate to the barrier to be permeated are used in the formulation. Such penetrants are generally known in the art, and
15 include, for example, for transmucosal administration, detergents, bile salts, and fusidic acid derivatives. Transmucosal administration can be accomplished through the use of nasal sprays or suppositories. For
20 transdermal administration, the active compounds are formulated into ointments, salves, gels, or creams as generally known in the art.

The compounds can also be prepared in the form of suppositories (e.g., with conventional suppository bases such as cocoa butter and other glycerides) or retention
25 enemas for rectal delivery.

In one embodiment, the active compounds are prepared with carriers that will protect the compound against rapid elimination from the body, such as a controlled release formulation, including implants and
30 microencapsulated delivery systems. Biodegradable, biocompatible polymers can be used, such as ethylene vinyl acetate, polyanhydrides, polyglycolic acid, collagen, polyorthoesters, and polylactic acid. Methods for preparation of such formulations will be apparent to
35 those skilled in the art. The materials can also be

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obtained commercially from Alza Corporation and Nova Pharmaceuticals, Inc. Liposomal suspensions (including liposomes targeted to infected cells with monoclonal antibodies to viral antigens) can also be used as
5 pharmaceutically acceptable carriers. These can be prepared according to methods known to those skilled in the art, for example, as described in U.S. Patent No. 4,522,811.

It is especially advantageous to formulate oral or
10 parenteral compositions in dosage unit form for ease of administration and uniformity of dosage. Dosage unit form as used herein refers to physically discrete units suited as unitary dosages for the subject to be treated; each unit containing a predetermined quantity of active
15 compound calculated to produce the desired therapeutic effect in association with the required pharmaceutical carrier. The specification for the dosage unit forms of the invention are dictated by and directly dependent on the unique characteristics of the active compound and the
20 particular therapeutic effect to be achieved, and the limitations inherent in the art of compounding such an active compound for the treatment of individuals.

The nucleic acid molecules of the invention can be inserted into vectors and used as gene therapy vectors.
25 Gene therapy vectors can be delivered to a subject by, for example, intravenous injection, local administration (U.S. Patent 5,328,470) or by stereotactic injection (see, e.g., Chen et al. (1994) *Proc. Natl. Acad. Sci. USA* 91:3054-3057). The pharmaceutical preparation of the
30 gene therapy vector can include the gene therapy vector in an acceptable diluent, or can comprise a slow release matrix in which the gene delivery vehicle is imbedded. Alternatively, where the complete gene delivery vector can be produced intact from recombinant cells, e.g.
35 retroviral vectors, the pharmaceutical preparation can

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include one or more cells which produce the gene delivery system.

The pharmaceutical compositions can be included in a container, pack, or dispenser together with
5 instructions for administration.

V. Uses and Methods of the Invention

The nucleic acid molecules, proteins, protein homologues, and antibodies described herein can be used in one or more of the following methods: a) screening
10 assays; b) detection assays (e.g., chromosomal mapping, tissue typing, forensic biology); c) predictive medicine (e.g., diagnostic assays, prognostic assays, monitoring clinical trials, and pharmacogenomics); and d) methods of treatment (e.g., therapeutic and prophylactic). A
15 Tango-77 protein interacts with other cellular proteins and can thus be used for regulation of inflammation. The polypeptides of the invention can be used in assays to determine biological activity. For example, they could be used in a panel of proteins for high-throughput
20 screening.

The isolated nucleic acid molecules of the invention can be used to express Tango-77 protein (e.g., via a recombinant expression vector in a host cell in gene therapy applications), to detect Tango-77 mRNA
25 (e.g., in a biological sample) or a genetic lesion in a Tango-77 gene, and to modulate Tango-77 activity. In addition, the Tango-77 proteins can be used to screen drugs or compounds which modulate the Tango-77 activity or expression as well as to treat disorders characterized
30 by insufficient or excessive production of Tango-77 protein or production of Tango-77 protein forms which have decreased or aberrant activity compared to Tango-77 wild type protein. In addition, the anti-Tango-77

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antibodies of the invention can be used to detect and isolate Tango-77 proteins and modulate Tango-77 activity.

This invention further pertains to novel agents identified by the above-described screening assays and
5 uses thereof for treatments as described herein.

A. Screening Assays

The invention provides a method (also referred to herein as a "screening assay") for identifying modulators, i.e., candidate or test compounds or agents
10 (e.g., peptides, peptidomimetics, small molecules or other drugs) which bind to Tango-77 proteins or have a stimulatory or inhibitory effect on, for example, Tango-77 expression or Tango-77 activity.

Examples of methods for the synthesis of molecular
15 libraries can be found in the art, for example in:
DeWitt et al. (1993) *Proc. Natl. Acad. Sci. USA* 90:6909;
Erb et al. (1994) *Proc. Natl. Acad. Sci. USA* 91:11422;
Zuckermann et al. (1994). *J. Med. Chem.* 37:2678; Cho et
al. (1993) *Science* 261:1303; Carrell et al. (1994) *Angew.*
20 *Chem. Int. Ed. Engl.* 33:2059; Carell et al. (1994) *Angew.*
Chem. Int. Ed. Engl. 33:2061; and Gallop et al. (1994) *J.*
Med. Chem. 37:1233.

Libraries of compounds may be presented in solution (e.g., Houghten (1992) *Bio/Techniques* 13:412-
25 421), or on beads (Lam (1991) *Nature* 354:82-84), chips (Fodor (1993) *Nature* 364:555-556), bacteria (U.S. Patent No. 5,223,409), spores (Patent Nos. 5,571,698; 5,403,484; and 5,223,409), plasmids (Cull et al. (1992) *Proc. Natl. Acad. Sci. USA* 89:1865-1869) or phage (Scott and Smith
30 (1990) *Science* 249:386-390; Devlin (1990) *Science* 249:404-406; Cwirla et al. (1990) *Proc. Natl. Acad. Sci. USA* 87:6378-6382; and Felici (1991) *J. Mol. Biol.* 222:301-310).

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In another embodiment, an assay is used to determine the ability of the test compound to modulate the activity of Tango-77 or a biologically active portion thereof, for example, by determining the ability of the

5 Tango-77 protein to bind to or interact with a Tango-77 target molecule. As used herein, a "target molecule" is a molecule with which a Tango-77 protein binds or interacts in nature, for example, a molecule on the surface of a cell. A Tango-77 target molecule can be a

10 non-Tango-77 molecule or a Tango-77 protein or polypeptide of the present invention. In one embodiment, a Tango-77 target molecule is a component of a signal transduction pathway, for example, Tango-77 may bind to a IL-1 receptor or another receptor thereby blocking the

15 receptor and inhibiting future signal transduction. Determining the ability of the Tango-77 protein to bind to or interact with a Tango-77 target molecule can be accomplished by one of the methods described above. In a preferred embodiment, determining the ability of the

20 Tango-77 protein to bind to or interact with a Tango-77 target molecule can be accomplished by determining the activity of the target molecule. For example, the activity of the target molecule can be determined by detecting induction of a cellular second messenger of the

25 target (e.g., intracellular Ca^{2+} , diacylglycerol, IP3, etc.), detecting catalytic/enzymatic activity of the target on an appropriate substrate, detecting the induction of a reporter gene (e.g., a Tango-77-responsive regulatory element operably linked to a nucleic acid

30 encoding a detectable marker, e.g. luciferase), or detecting a cellular response, for example, inflammation.

In yet another embodiment, an assay of the present invention is a cell-free assay comprising contacting a Tango-77 protein or biologically active portion thereof

35 with a test compound and determining the ability of the

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test compound to bind to the Tango-77 protein or biologically active portion thereof. Binding of the test compound to the Tango-77 protein can be determined either directly or indirectly as described above. In a preferred embodiment, the assay includes contacting the Tango-77 protein or biologically active portion thereof with a known compound which binds Tango-77 to form an assay mixture, contacting the assay mixture with a test compound, and determining the ability of the test compound to interact with a Tango-77 protein, wherein determining the ability of the test compound to interact with a Tango-77 protein comprises determining the ability of the test compound to preferentially bind to Tango-77 or biologically active portion thereof as compared to the known compound.

In another embodiment, an assay is a cell-free assay comprising contacting Tango-77 protein or biologically active portion thereof with a test compound and determining the ability of the test compound to modulate (e.g., stimulate or inhibit) the activity of the Tango-77 protein or biologically active portion thereof. Determining the ability of the test compound to modulate the activity of Tango-77 can be accomplished, for example, by determining the ability of the Tango-77 protein to bind to a Tango-77 target molecule by one of the methods described above for determining direct binding. In an alternative embodiment, determining the ability of the test compound to modulate the activity of Tango-77 can be accomplished by determining the ability of the Tango-77 protein to further modulate a Tango-77 target molecule. For example, the catalytic/enzymatic activity of the target molecule on an appropriate substrate can be determined as previously described.

In yet another embodiment, the cell-free assay comprises contacting the Tango-77 protein or biologically

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active portion thereof with a known compound which binds Tango-77 to form an assay mixture, contacting the assay mixture with a test compound, and determining the ability of the test compound to interact with a Tango-77 protein, wherein determining the ability of the test compound to interact with a Tango-77 protein comprises determining the ability of the Tango-77 protein to preferentially bind to or modulate the activity of a Tango-77 target molecule.

10 It is possible that membrane-bound forms of Tango-77 exist. The cell-free assays of the present invention are amenable to use of both the forms Tango-77. In the case of cell-free assays comprising a membrane-bound form of Tango-77, it may be desirable to utilize a
15 solubilizing agent such that the membrane-bound form of Tango-77 is maintained in solution. Examples of such solubilizing agents include non-ionic detergents such as n-octylglucoside, n-dodecylglucoside, n-dodecylmaltoside, octanoyl-N-methylglucamide, decanoyl-N-methylglucamide,
20 Triton® X-100, Triton® X-114, Thesit®, Isotridecypoly(ethylene glycol ether)n, 3-[(3-cholamidopropyl)dimethylamminio]-1-propane sulfonate (CHAPS), 3-[(3-cholamidopropyl)dimethylamminio]-2-hydroxy-1-propane sulfonate (CHAPSO), or N-dodecyl=N,N-
25 dimethyl-3-ammonio-1-propane sulfonate.

In more than one embodiment of the above assay methods of the present invention, it may be desirable to immobilize either Tango-77 or its target molecule to facilitate separation of complexed from uncomplexed forms
30 of one or both of the proteins, as well as to accommodate automation of the assay. Binding of a test compound to Tango-77, or interaction of Tango-77 with a target molecule in the presence and absence of a candidate compound, can be accomplished in any vessel suitable for
35 containing the reactants. Examples of such vessels

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include microtitre plates, test tubes, and micro-centrifuge tubes. In one embodiment, a fusion protein can be provided which adds a domain that allows one or both of the proteins to be bound to a matrix. For
5 example, glutathione-S-transferase/ Tango-77 fusion proteins or glutathione-S-transferase/target fusion proteins can be adsorbed onto glutathione sepharose beads (Sigma Chemical Co.; St. Louis, MO) or glutathione derivatized microtitre plates, which are then combined
10 with the test compound or the test compound and either the non-adsorbed target protein or Tango-77 protein, and the mixture incubated under conditions conducive to complex formation (e.g., at physiological conditions for salt and pH). Following incubation, the beads or
15 microtitre plate wells are washed to remove any unbound components and complex formation is measured either directly or indirectly, for example, as described above. Alternatively, the complexes can be dissociated from the matrix, and the level of Tango-77 binding or activity
20 determined using standard techniques.

Other techniques for immobilizing proteins on matrices can also be used in the screening assays of the invention. For example, either Tango-77 or its target molecule can be immobilized utilizing conjugation of
25 biotin and streptavidin. Biotinylated Tango-77 or target molecules can be prepared from biotin-NHS (N-hydroxy-succinimide) using techniques well known in the art (e.g., biotinylation kit, Pierce Chemicals; Rockford, IL), and immobilized in the wells of streptavidin-coated
30 96 well plates (Pierce Chemical). Alternatively, antibodies reactive with Tango-77 or target molecules but which do not interfere with binding of the Tango-77 protein to its target molecule can be derivatized to the wells of the plate, and unbound target or Tango-77
35 trapped in the wells by antibody conjugation. Methods

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for detecting such complexes, in addition to those described above for the GST-immobilized complexes, include immunodetection of complexes using antibodies reactive with the Tango-77 or target molecule, as well as
5 enzyme-linked assays which rely on detecting an enzymatic activity associated with the Tango-77 or target molecule.

In another embodiment, modulators of Tango-77 expression are identified in a method in which a cell is contacted with a candidate compound and the expression of
10 Tango-77 mRNA or protein in the cell is determined. The level of expression of Tango-77 mRNA or protein in the presence of the candidate compound is compared to the level of expression of Tango-77 mRNA or protein in the absence of the candidate compound. The candidate
15 compound can then be identified as a modulator of Tango-77 expression based on this comparison. For example, when expression of Tango-77 mRNA or protein is greater (statistically significantly greater) in the presence of the candidate compound than in its absence,
20 the candidate compound is identified as a stimulator of Tango-77 mRNA or protein expression. Alternatively, when expression of Tango-77 mRNA or protein is less (statistically significantly less) in the presence of the candidate compound than in its absence, the candidate
25 compound is identified as an inhibitor of Tango-77 mRNA or protein expression. The level of Tango-77 mRNA or protein expression in the cells can be determined by methods described herein for detecting Tango-77 mRNA or protein.

30 In yet another aspect of the invention, the Tango-77 proteins can be used as "bait proteins" in a two-hybrid assay or three hybrid assay (see, e.g., U.S. Patent No. 5,283,317; Zervos et al. (1993) *Cell* 72:223-232; Madura et al. (1993) *J. Biol. Chem.* 268:12046-12054;
35 Bartel et al. (1993) *Bio/Techniques* 14:920-924; Iwabuchi

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et al. (1993) *Oncogene* 8:1693-1696; and PCT Publication No. WO 94/10300), to identify other proteins, which bind to or interact with Tango-77 ("Tango-77-binding proteins" or "Tango-77-bp") and modulate Tango-77 activity. Such
5 Tango-77-binding proteins are also likely to be involved in the propagation of signals by the Tango-77 proteins as, for example, upstream or downstream elements of the Tango-77 pathway.

The two-hybrid system is based on the modular
10 nature of most transcription factors, which consist of separable DNA-binding and activation domains. Briefly, the assay utilizes two different DNA constructs. In one construct, the gene that codes for Tango-77 is fused to a gene encoding the DNA binding domain of a known
15 transcription factor (e.g., GAL-4). In the other construct, a DNA sequence, from a library of DNA sequences, that encodes an unidentified protein ("prey" or "sample") is fused to a gene that codes for the activation domain of the known transcription factor. If
20 the "bait" and the "prey" proteins are able to interact, *in vivo*, forming an Tango-77-dependent complex, the DNA-binding and activation domains of the transcription factor are brought into close proximity. This proximity allows transcription of a reporter gene (e.g., LacZ)
25 which is operably linked to a transcriptional regulatory site responsive to the transcription factor. Expression of the reporter gene can be detected and cell colonies containing the functional transcription factor can be isolated and used to obtain the cloned gene which encodes
30 the protein which interacts with Tango-77.

This invention further pertains to novel agents identified by the above-described screening assays and uses thereof for treatments as described herein.

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B. Detection Assays

Portions or fragments of the cDNA sequence identified herein (and the corresponding complete gene sequences) can be used in numerous ways as polynucleotide reagents. For example, the sequence can be used to: (i) map the respective gene on a chromosome and, thus, locate gene regions associated with genetic disease; (ii) identify an individual from a minute biological sample (tissue typing); and (iii) aid in forensic identification of a biological sample. These applications are described in the subsections below.

1. Chromosome Mapping

Once the sequence (or a portion of the sequence) of a gene has been isolated, this sequence can be used to map the location of the gene on a chromosome. Accordingly, Tango-77 nucleic acid molecules described herein or fragments thereof, can be used to map the location of the Tango-77 gene(s) on a chromosome. The mapping of the Tango-77 sequences to chromosomes is an important first step in correlating these sequences with genes associated with disease.

Briefly, a Tango-77 gene can be mapped to chromosomes by preparing PCR primers (preferably 15-25 bp in length) from the Tango-77 sequences. Computer analysis of Tango-77 sequences can be used to rapidly select primers that do not span more than one exon in the genomic DNA, thus complicating the amplification process. These primers can then be used for PCR screening of somatic cell hybrids containing individual human chromosomes. Only those hybrids containing the human gene corresponding to the Tango-77 sequences will yield an amplified fragment.

Somatic cell hybrids are prepared by fusing somatic cells from different mammals (e.g., human and

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mouse cells). As hybrids of human and mouse cells grow and divide, they gradually lose human chromosomes in random order, but retain the mouse chromosomes. By using media in which mouse cells cannot grow (because they lack a particular enzyme) but in which human cells can, the one human chromosome that contains the gene encoding the needed enzyme, will be retained. By using various media, panels of hybrid cell lines can be established. Each cell line in a panel contains either a single human chromosome or a small number of human chromosomes, and a full set of mouse chromosomes, allowing easy mapping of individual genes to specific human chromosomes. (D'Eustachio et al. (1983) *Science* 220:919-924). Somatic cell hybrids containing only fragments of human chromosomes can also be produced by using human chromosomes with translocations and deletions.

PCR mapping of somatic cell hybrids is a rapid procedure for assigning a particular sequence to a particular chromosome. Three or more sequences can be assigned per day using a single thermal cycler. Using the Tango-77 sequences to design oligonucleotide primers, sublocalization can be achieved with panels of fragments from specific chromosomes. Other mapping strategies which can similarly be used to map a Tango-77 sequence to its chromosome include *in situ* hybridization (described in Fan et al. (1990) *Proc. Natl. Acad. Sci. USA* 87:6223-27), pre-screening with labeled flow-sorted chromosomes, and pre-selection by hybridization to chromosome specific cDNA libraries.

Fluorescence *in situ* hybridization (FISH) of a DNA sequence to a metaphase chromosomal spread can further be used to provide a precise chromosomal location in one step. Chromosome spreads can be made using cells whose division has been blocked in metaphase by a chemical, e.g., colcemid that disrupts the mitotic spindle. The

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chromosomes can be treated briefly with trypsin, and then stained with Giemsa. A pattern of light and dark bands develops on each chromosome, so that the chromosomes can be identified individually. The FISH technique can be
5 used with a DNA sequence as short as 500 or 600 bases. However, clones larger than 1,000 bases have a higher likelihood of binding to a unique chromosomal location with sufficient signal intensity for simple detection. Preferably 1,000 bases, and more preferably 2,000 bases
10 will suffice to get good results at a reasonable amount of time. For a review of this technique, see Verma et al. (Human Chromosomes: A Manual of Basic Techniques (Pergamon Press, New York, 1988)).

Reagents for chromosome mapping can be used
15 individually to mark a single chromosome or a single site on that chromosome, or panels of reagents can be used for marking multiple sites and/or multiple chromosomes. Reagents corresponding to noncoding regions of the genes actually are preferred for mapping purposes. Coding
20 sequences are more likely to be conserved within gene families, thus increasing the chance of cross hybridizations during chromosomal mapping.

Once a sequence has been mapped to a precise chromosomal location, the physical position of the
25 sequence on the chromosome can be correlated with genetic map data. (Such data are found, for example, in V. McKusick, Mendelian Inheritance in Man, available on-line through Johns Hopkins University Welch Medical Library). The relationship between genes and disease, mapped to the
30 same chromosomal region, can then be identified through linkage analysis (co-inheritance of physically adjacent genes), described in, e.g., Egeland et al. (1987) Nature 325:783-787.

Moreover, differences in the DNA sequences between
35 individuals affected and unaffected with a disease

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associated with the Tango-77 gene can be determined. If a mutation is observed in some or all of the affected individuals but not in any unaffected individuals, then the mutation is likely to be the causative agent of the particular disease. Comparison of affected and unaffected individuals generally involves first looking for structural alterations in the chromosomes such as deletions or translocations that are visible from chromosome spreads or detectable using PCR based on that DNA sequence. Ultimately, complete sequencing of genes from several individuals can be performed to confirm the presence of a mutation and to distinguish mutations from polymorphisms.

2. Tissue Typing

The Tango-77 sequences of the present invention can also be used to identify individuals from minute biological samples. The United States military, for example, is considering the use of restriction fragment length polymorphism (RFLP) for identification of its personnel. In this technique, an individual's genomic DNA is digested with one or more restriction enzymes, and probed on a Southern blot to yield unique bands for identification. This method does not suffer from the current limitations of "Dog Tags" which can be lost, switched, or stolen, making positive identification difficult. The sequences of the present invention are useful as additional DNA markers for RFLP (described in U.S. Patent 5,272,057).

Furthermore, the sequences of the present invention can be used to provide an alternative technique which determines the actual base-by-base DNA sequence of selected portions of an individual's genome. Thus, the Tango-77 sequences described herein can be used to prepare two PCR primers from the 5' and 3' ends of the

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sequences. These primers can then be used to amplify an individual's DNA and subsequently sequence it.

Panels of corresponding DNA sequences from individuals, prepared in this manner, can provide unique individual identifications, as each individual will have a unique set of such DNA sequences due to allelic differences. The sequences of the present invention can be used to obtain such identification sequences from individuals and from tissue. The Tango-77 sequences of the invention uniquely represent portions of the human genome. Allelic variation occurs to some degree in the coding regions of these sequences, and to a greater degree in the noncoding regions. It is estimated that allelic variation between individual humans occurs with a frequency of about once per each 500 bases. Each of the sequences described herein can, to some degree, be used as a standard against which DNA from an individual can be compared for identification purposes. Because greater numbers of polymorphisms occur in the noncoding regions, fewer sequences are necessary to differentiate individuals. The noncoding sequences of SEQ ID NO:1 can comfortably provide positive individual identification with a panel of perhaps 10 to 1,000 primers which each yield a noncoding amplified sequence of 100 bases. If predicted coding sequences, such as those in SEQ ID NO:3, SEQ ID NO:6, or SEQ ID NO:10 are used, a more appropriate number of primers for positive individual identification would be 500-2,000.

If a panel of reagents from Tango-77 sequences described herein is used to generate a unique identification database for an individual, those same reagents can later be used to identify tissue from that individual. Using the unique identification database, positive identification of the individual, living or dead, can be made from extremely small tissue samples.

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3. Use of Partial Tango-77 Sequences in Forensic Biology

DNA-based identification techniques can also be used in forensic biology. Forensic biology is a scientific field employing genetic typing of biological evidence found at a crime scene as a means for positively identifying, for example, a perpetrator of a crime. To make such an identification, PCR technology can be used to amplify DNA sequences taken from very small biological samples such as tissues, e.g., hair or skin, or body fluids, e.g., blood, saliva, or semen found at a crime scene. The amplified sequence can then be compared to a standard, thereby allowing identification of the origin of the biological sample.

The sequences of the present invention can be used to provide polynucleotide reagents, e.g., PCR primers, targeted to specific loci in the human genome, which can enhance the reliability of DNA-based forensic identifications by, for example, providing another "identification marker" (i.e. another DNA sequence that is unique to a particular individual). As mentioned above, actual base sequence information can be used for identification as an accurate alternative to patterns formed by restriction enzyme generated fragments. Sequences targeted to noncoding regions of SEQ ID NO:1 are particularly appropriate for this use as greater numbers of polymorphisms occur in the noncoding regions, making it easier to differentiate individuals using this technique. Examples of polynucleotide reagents include the Tango-77 sequences or portions thereof, e.g., fragments derived from the noncoding regions of SEQ ID NO:1 having a length of at least 20 or 30 bases.

The Tango-77 sequences described herein can further be used to provide polynucleotide reagents, e.g., labeled or labelable probes which can be used in, for

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example, an in situ hybridization technique, to identify a specific tissue, e.g., brain tissue. This can be very useful in cases where a forensic pathologist is presented with a tissue of unknown origin. Panels of such Tango-77 probes can be used to identify tissue by species and/or by organ type.

In a similar fashion, these reagents, e.g., Tango-77 primers or probes can be used to screen tissue culture for contamination (i.e., screen for the presence of a mixture of different types of cells in a culture).

C. Predictive Medicine

The present invention also pertains to the field of predictive medicine in which diagnostic assays, prognostic assays, pharmacogenomics, and monitoring clinical trials are used for prognostic (predictive) purposes to thereby treat an individual prophylactically. Accordingly, one aspect of the present invention relates to diagnostic assays for determining Tango-77 protein and/or nucleic acid expression as well as Tango-77 activity, in the context of a biological sample (e.g., blood, serum, cells, tissue) to thereby determine whether an individual is afflicted with a disease or disorder, or is at risk of developing a disorder, associated with aberrant Tango-77 expression or activity. The invention also provides for prognostic (or predictive) assays for determining whether an individual is at risk of developing a disorder associated with Tango-77 protein, nucleic acid expression or activity. For example, mutations in a Tango-77 gene can be assayed in a biological sample. Such assays can be used for prognostic or predictive purpose to thereby prophylactically treat an individual prior to the onset of a disorder characterized by or associated with Tango-77 protein, nucleic acid expression or activity.

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Another aspect of the invention provides methods for determining Tango-77 protein, nucleic acid expression or Tango-77 activity in an individual to thereby select appropriate therapeutic or prophylactic agents for that individual (referred to herein as "pharmacogenomics"). Pharmacogenomics allows for the selection of agents (e.g., drugs) for therapeutic or prophylactic treatment of an individual based on the genotype of the individual (e.g., the genotype of the individual examined to determine the ability of the individual to respond to a particular agent.)

Yet another aspect of the invention pertains to monitoring the influence of agents (e.g., drugs or other compounds) on the expression or activity of Tango-77 in clinical trials.

These and other agents are described in further detail in the following sections.

1. Diagnostic Assays

An exemplary method for detecting the presence or absence of Tango-77 in a biological sample involves obtaining a biological sample from a test subject and contacting the biological sample with a compound or an agent capable of detecting Tango-77 protein or nucleic acid (e.g., mRNA, genomic DNA) that encodes Tango-77 protein such that the presence of Tango-77 is detected in the biological sample. A preferred agent for detecting Tango-77 mRNA or genomic DNA is a labeled nucleic acid probe capable of hybridizing to Tango-77 mRNA or genomic DNA. The nucleic acid probe can be, for example, a full-length Tango-77 nucleic acid, such as the nucleic acid of SEQ ID NO: 1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10 or a portion thereof, such as an oligonucleotide of at least 15, 30, 50, 100, 250 or 500 nucleotides in length and sufficient to specifically hybridize under stringent

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conditions to Tango-77 mRNA or genomic DNA. Other suitable probes for use in the diagnostic assays of the invention are described herein.

A preferred agent for detecting Tango-77 protein is an antibody capable of binding to Tango-77 protein, preferably an antibody with a detectable label. Antibodies can be polyclonal, or more preferably, monoclonal. An intact antibody, or a fragment thereof (e.g., Fab or F(ab')₂) can be used. The term "labeled", with regard to the probe or antibody, is intended to encompass direct labeling of the probe or antibody by coupling (i.e., physically linking) a detectable substance to the probe or antibody, as well as indirect labeling of the probe or antibody by reactivity with another reagent that is directly labeled. Examples of indirect labeling include detection of a primary antibody using a fluorescently labeled secondary antibody and end-labeling of a DNA probe with biotin such that it can be detected with fluorescently labeled streptavidin. The term "biological sample" is intended to include tissues, cells and biological fluids isolated from a subject, as well as tissues, cells and fluids present within a subject. That is, the detection method of the invention can be used to detect Tango-77 mRNA, protein, or genomic DNA in a biological sample in vitro as well as in vivo. For example, in vitro techniques for detection of Tango-77 mRNA include Northern hybridizations and in situ hybridizations. In vitro techniques for detection of Tango-77 protein include enzyme linked immunosorbent assays (ELISAs), Western blots, immunoprecipitations and immunofluorescence. In vitro techniques for detection of Tango-77 genomic DNA include Southern hybridizations. Furthermore, in vivo techniques for detection of Tango-77 protein include introducing into a subject a labeled anti-Tango-77 antibody. For example, the antibody can be

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labeled with a radioactive marker whose presence and location in a subject can be detected by standard imaging techniques.

In one embodiment, the biological sample contains
5 protein molecules from the test subject. Alternatively, the biological sample can contain mRNA molecules from the test subject or genomic DNA molecules from the test subject. A preferred biological sample is a peripheral blood leukocyte sample isolated by conventional means
10 from a subject.

In another embodiment, the methods further involve obtaining a control biological sample from a control subject, contacting the control sample with a compound or agent capable of detecting Tango-77 protein, mRNA, or
15 genomic DNA, such that the presence of Tango-77 protein, mRNA or genomic DNA is detected in the biological sample, and comparing the presence of Tango-77 protein, mRNA or genomic DNA in the control sample with the presence of Tango-77 protein, mRNA or genomic DNA in the test sample.

20 The invention also encompasses kits for detecting the presence of Tango-77 in a biological sample (a test sample). Such kits can be used to determine if a subject is suffering from or is at increased risk of developing a disorder associated with aberrant expression of Tango-77
25 (e.g., an immunological disorder). For example, the kit can comprise a labeled compound or agent capable of detecting Tango-77 protein or mRNA in a biological sample and means for determining the amount of Tango-77 in the sample (e.g., an anti-Tango-77 antibody or an
30 oligonucleotide probe which binds to DNA encoding Tango-77, e.g., SEQ ID NO:1 or SEQ ID NO:3 or SEQ ID NO:6, or SEQ ID NO:10). Kits may also include instruction for observing that the tested subject is suffering from or is at risk of developing a disorder
35 associated with aberrant expression of Tango-77 if the

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amount of Tango-77 protein or mRNA is above or below a normal level.

For antibody-based kits, the kit may comprise, for example: (1) a first antibody (e.g., attached to a solid support) which binds to Tango-77 protein; and, optionally (2) a second, different antibody which binds to Tango-77 protein or the first antibody and is conjugated to a detectable agent.

For oligonucleotide-based kits, the kit may comprise, for example: (1) an oligonucleotide, e.g., a detectably labelled oligonucleotide, which hybridizes to a Tango-77 nucleic acid sequence or (2) a pair of primers useful for amplifying a Tango-77 nucleic acid molecule;

The kit may also comprise, e.g., a buffering agent, a preservative, or a protein stabilizing agent. The kit may also comprise components necessary for detecting the detectable agent (e.g., an enzyme or a substrate). The kit may also contain a control sample or a series of control samples which can be assayed and compared to the test sample contained. Each component of the kit is usually enclosed within an individual container and all of the various containers are within a single package along with instructions for observing whether the tested subject is suffering from or is at risk of developing a disorder associated with aberrant expression of Tango-77.

2. Prognostic Assays

The methods described herein can furthermore be utilized as diagnostic or prognostic assays to identify subjects having or at risk of developing a disease or disorder associated with aberrant Tango-77 expression or activity. For example, the assays described herein, such as the preceding diagnostic assays or the following assays, can be utilized to identify a subject having or

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at risk of developing a disorder associated with aberrant expression or activity. Thus, the present invention provides a method in which a test sample is obtained from a subject and Tango-77 protein or nucleic acid (e.g., mRNA, genomic DNA) is detected, wherein the presence of Tango-77 protein or nucleic acid is diagnostic for a subject having or at risk of developing a disease or disorder associated with aberrant Tango-77 expression or activity. As used herein, a "test sample" refers to a biological sample obtained from a subject of interest. For example, a test sample can be a biological fluid (e.g., serum), cell sample, or tissue.

Furthermore, the prognostic assays described herein can be used to determine whether a subject can be administered an agent (e.g., an agonist, antagonist, peptidomimetic, protein, peptide, nucleic acid, small molecule, or other drug candidate) to treat a disease or disorder associated with aberrant Tango-77 expression or activity. For example, such methods can be used to determine whether a subject can be effectively treated with a specific agent or class of agents (e.g., agents of a type which decrease Tango-77 activity). Thus, the present invention provides methods for determining whether a subject can be effectively treated with an agent for a disorder associated with aberrant Tango-77 expression or activity in which a test sample is obtained and Tango-77 protein or nucleic acid is detected (e.g., wherein the presence of Tango-77 protein or nucleic acid is diagnostic for a subject that can be administered the agent to treat a disorder associated with aberrant Tango-77 expression or activity).

The methods of the invention can also be used to detect genetic lesions or mutations in a Tango-77 gene, thereby determining if a subject with the lesioned gene is at risk for a disorder characterized by aberrant

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inflammation. In preferred embodiments, the methods include detecting, in a sample of cells from the subject, the presence or absence of a genetic lesion or mutation characterized by at least one of an alteration affecting
5 the integrity of a gene encoding a Tango-77-protein, or the mis-expression of the Tango-77 gene. For example, such genetic lesions or mutations can be detected by ascertaining the existence of at least one of: 1) a deletion of one or more nucleotides from a Tango-77 gene;
10 2) an addition of one or more nucleotides to a Tango-77 gene; 3) a substitution of one or more nucleotides of a Tango-77 gene; 4) a chromosomal rearrangement of a Tango-77 gene; 5) an alteration in the level of a messenger RNA transcript of a Tango-77 gene; 6) an
15 aberrant modification of a Tango-77 gene, such as of the methylation pattern of the genomic DNA; 7) the presence of a non-wild type splicing pattern of a messenger RNA transcript of a Tango-77 gene; 8) a non-wild type level of a Tango-77-protein; 9) an allelic loss of a Tango-77
20 gene, and 10) an inappropriate post-translational modification of a Tango-77-protein. As described herein, there are a large number of assay techniques known in the art which can be used for detecting lesions or mutations in a Tango-77 gene. A preferred biological sample is a
25 peripheral blood leukocyte sample isolated by conventional means from a subject.

In certain embodiments, detection of the lesion involves the use of a probe/primer in a polymerase chain reaction (PCR) (see, e.g., U.S. Patent Nos. 4,683,195 and
30 4,683,202), such as anchor PCR or RACE PCR, or, alternatively, in a ligation chain reaction (LCR) (see, e.g., Landegran et al. (1988) *Science* 241:1077-1080; and Nakazawa et al. (1994) *Proc. Natl. Acad. Sci. USA* 91:360-364), the latter of which can be particularly useful for
35 detecting point mutations in the Tango-77-gene (see,

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e.g., Abravaya et al. (1995) *Nucleic Acids Res.* 23:675-682). This method can include the steps of collecting a sample of cells from a patient, isolating nucleic acid (e.g., genomic, mRNA or both) from the cells of the sample, contacting the nucleic acid sample with one or more primers which specifically hybridize to a Tango-77 gene under conditions such that hybridization and amplification of the Tango-77-gene (if present) occurs, and detecting the presence or absence of an amplification product, or detecting the size of the amplification product and comparing the length to a control sample. It is anticipated that PCR and/or LCR may be desirable to use as a preliminary amplification step in conjunction with any of the techniques used for detecting mutations described herein.

Alternative amplification methods include: self sustained sequence replication (Guatelli et al. (1990) *Proc. Natl. Acad. Sci. USA* 87:1874-1878), transcriptional amplification system (Kwoh, et al. (1989) *Proc. Natl. Acad. Sci. USA* 86:1173-1177), Q-Beta Replicase (Lizardi et al. (1988) *Bio/Technology* 6:1197), or any other nucleic acid amplification method, followed by the detection of the amplified molecules using techniques well known to those of skill in the art. These detection schemes are especially useful for the detection of nucleic acid molecules if such molecules are present in very low numbers.

In an alternative embodiment, mutations in a Tango-77 gene from a sample cell can be identified by alterations in restriction enzyme cleavage patterns. For example, sample and control DNA is isolated, amplified (optionally), digested with one or more restriction endonucleases, and fragment length sizes are determined by gel electrophoresis and compared. Differences in fragment length sizes between sample and control DNA

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indicates mutations in the sample DNA. Moreover, the use of sequence specific ribozymes (see, e.g., U.S. Patent No. 5,498,531) can be used to score for the presence of specific mutations by development or loss of a ribozyme cleavage site.

In other embodiments, genetic mutations in Tango-77 can be identified by hybridizing a sample and control nucleic acids, e.g., DNA or RNA, to high density arrays containing hundreds or thousands of oligonucleotide probes (Cronin et al. (1996) *Human Mutation* 7:244-255; Kozal et al. (1996) *Nature Medicine* 2:753-759). For example, genetic mutations in Tango-77 can be identified in two-dimensional arrays containing light-generated DNA probes as described in Cronin et al. supra. Briefly, a first hybridization array of probes can be used to scan through long stretches of DNA in a sample and control to identify base changes between the sequences by making linear arrays of sequential overlapping probes. This step allows the identification of point mutations. This step is followed by a second hybridization array that allows the characterization of specific mutations by using smaller, specialized probe arrays complementary to all variants or mutations detected. Each mutation array is composed of parallel probe sets, one complementary to the wild-type gene and the other complementary to the mutant gene.

In yet another embodiment, any of a variety of sequencing reactions known in the art can be used to directly sequence the Tango-77 gene and detect mutations by comparing the sequence of the sample Tango-77 with the corresponding wild-type (control) sequence. Examples of sequencing reactions include those based on techniques developed by Maxim and Gilbert ((1977) *Proc. Natl. Acad. Sci. USA* 74:560) or Sanger ((1977) *Proc. Natl. Acad. Sci. USA* 74:5463). It is also contemplated that any of a

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variety of automated sequencing procedures can be utilized when performing the diagnostic assays ((1995) *Bio/Techniques* 19:448), including sequencing by mass spectrometry (see, e.g., PCT Publication No. WO 94/16101; 5 Cohen et al. (1996) *Adv. Chromatogr.* 36:127-162; and Griffin et al. (1993) *Appl. Biochem. Biotechnol.* 38:147-159).

Other methods for detecting mutations in the Tango-77 gene include methods in which protection from 10 cleavage agents is used to detect mismatched bases in RNA/RNA or RNA/DNA heteroduplexes (Myers et al. (1985) *Science* 230:1242). In general, the technique of "mismatch cleavage" entails providing heteroduplexes formed by hybridizing (labeled) RNA or DNA containing the 15 wild-type Tango-77 sequence with potentially mutant RNA or DNA obtained from a tissue sample. The double-stranded duplexes are treated with an agent which cleaves single-stranded regions of the duplex such as which will exist due to basepair mismatches between the control and 20 sample strands. RNA/DNA duplexes can be treated with RNase to digest mismatched regions, and DNA/DNA hybrids can be treated with S1 nuclease to digest mismatched regions. In other embodiments, either DNA/DNA or RNA/DNA duplexes can be treated with hydroxylamine or osmium 25 tetroxide and with piperidine in order to digest mismatched regions. After digestion of the mismatched regions, the resulting material is then separated by size on denaturing polyacrylamide gels to determine the site of mutation. See, e.g., Cotton et al. (1988) *Proc. Natl. 30 Acad. Sci. USA* 85:4397; Saleeba et al. (1992) *Methods Enzymol.* 217:286-295. In a preferred embodiment, the control DNA or RNA can be labeled for detection.

In still another embodiment, the mismatch cleavage reaction employs one or more proteins that recognize 35 mismatched base pairs in double-stranded DNA (so called

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"DNA mismatch repair" enzymes) in defined systems for detecting and mapping point mutations in Tango-77 cDNAs obtained from samples of cells. For example, the mutY enzyme of *E. coli* cleaves A at G/A mismatches and the
5 thymidine DNA glycosylase from HeLa cells cleaves T at G/T mismatches (Hsu et al. (1994) *Carcinogenesis* 15:1657-1662). According to an exemplary embodiment, a probe based on a Tango-77 sequence, e.g., a wild-type Tango-77
10 sequence, is hybridized to a cDNA or other DNA product from a test cell(s). The duplex is treated with a DNA mismatch repair enzyme, and the cleavage products, if any, can be detected from electrophoresis protocols or the like. See, e.g., U.S. Patent No. 5,459,039.

In other embodiments, alterations in
15 electrophoretic mobility will be used to identify mutations in Tango-77 genes. For example, single strand conformation polymorphism (SSCP) may be used to detect differences in electrophoretic mobility between mutant and wild type nucleic acids (Orita et al. (1989) *Proc.*
20 *Natl. Acad. Sci. USA* 86:2766; see also Cotton (1993) *Mutat. Res.* 285:125-144; Hayashi (1992) *Genet Anal Tech Appl* 9:73-79). Single-stranded DNA fragments of sample and control Tango-77 nucleic acids will be denatured and allowed to renature. The secondary structure of single-
25 stranded nucleic acids varies according to sequence, and the resulting alteration in electrophoretic mobility enables the detection of even a single base change. The DNA fragments may be labeled or detected with labeled probes. The sensitivity of the assay may be enhanced by
30 using RNA (rather than DNA), in which the secondary structure is more sensitive to a change in sequence. In a preferred embodiment, the subject method utilizes heteroduplex analysis to separate double stranded heteroduplex molecules on the basis of changes in

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electrophoretic mobility (Keen et al. (1991) *Trends Genet* 7:5).

In yet another embodiment, the movement of mutant or wild-type fragments in polyacrylamide gels containing
5 a gradient of denaturant is assayed using denaturing gradient gel electrophoresis (DGGE) (Myers et al. (1985) *Nature* 313:495). When DGGE is used as the method of analysis, DNA will be modified to insure that it does not completely denature, for example by adding a GC clamp of
10 approximately 40 bp of high-melting GC-rich DNA by PCR. In a further embodiment, a temperature gradient is used in place of a denaturing gradient to identify differences in the mobility of control and sample DNA (Rosenbaum and Reissner (1987) *Biophys. Chem.* 265:12753).

15 Examples of other techniques for detecting point mutations include, but are not limited to, selective oligonucleotide hybridization, selective amplification, or selective primer extension. For example, oligonucleotide primers may be prepared in which the
20 known mutation is placed centrally and then hybridized to target DNA under conditions which permit hybridization only if a perfect match is found (Saiki et al. (1986) *Nature* 324:163); Saiki et al. (1989) *Proc. Natl. Acad. Sci. USA* 86:6230). Such allele specific oligonucleotides
25 are hybridized to PCR amplified target DNA or a number of different mutations when the oligonucleotides are attached to the hybridizing membrane and hybridized with labeled target DNA.

Alternatively, allele specific amplification
30 technology which depends on selective PCR amplification may be used in conjunction with the instant invention. Oligonucleotides used as primers for specific amplification may carry the mutation of interest in the center of the molecule (so that amplification depends on
35 differential hybridization) (Gibbs et al. (1989) *Nucleic*

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Acids Res. 17:2437-2448) or at the extreme 3' end of one primer where, under appropriate conditions, mismatch can prevent or reduce polymerase extension (Prossner (1993) Tibtech 11:238). In addition, it may be desirable to
5 introduce a novel restriction site in the region of the mutation to create cleavage-based detection (Gasparini et al. (1992) Mol. Cell Probes 6:1). It is anticipated that in certain embodiments amplification may also be performed using Taq ligase for amplification (Barany
10 (1991) Proc. Natl. Acad. Sci USA 88:189). In such cases, ligation will occur only if there is a perfect match at the 3' end of the 5' sequence making it possible to detect the presence of a known mutation at a specific site by looking for the presence or absence of
15 amplification.

The methods described herein may be performed, for example, by utilizing pre-packaged diagnostic kits comprising at least one probe nucleic acid or antibody reagent described herein, which may be conveniently used,
20 e.g., in clinical settings to diagnose patients exhibiting symptoms or family history of a disease or illness involving a Tango-77 gene.

Furthermore, any cell type or tissue, preferably peripheral blood leukocytes, in which Tango-77 is
25 expressed may be utilized in the prognostic assays described herein.

3. Pharmacogenomics

Agents, or modulators which have a stimulatory or
30 inhibitory effect on Tango-77 activity (e.g., Tango-77 gene expression) as identified by a screening assay described herein can be administered to individuals to treat (prophylactically or therapeutically) disorders (e.g., acute or chronic inflammation and asthma)
35 associated with aberrant Tango-77 activity. In

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conjunction with such treatment, the pharmacogenomics (i.e., the study of the relationship between an individual's genotype and that individual's response to a foreign compound or drug) of the individual may be considered. Differences in metabolism of therapeutics can lead to severe toxicity or therapeutic failure by altering the relation between dose and blood concentration of the pharmacologically active drug. Thus, the pharmacogenomics of the individual permits the selection of effective agents (e.g., drugs) for prophylactic or therapeutic treatments based on a consideration of the individual's genotype. Such pharmacogenomics can further be used to determine appropriate dosages and therapeutic regimens. Accordingly, the activity of Tango-77 protein, expression of Tango-77 nucleic acid, or mutation content of Tango-77 genes in an individual can be determined to thereby select appropriate agent(s) for therapeutic or prophylactic treatment of the individual.

Pharmacogenomics deals with clinically significant hereditary variations in the response to drugs due to altered drug disposition and abnormal action in affected persons. See, e.g., Linder (1997) *Clin. Chem.* 43(2):254-266. In general, two types of pharmacogenetic conditions can be differentiated. Genetic conditions transmitted as a single factor altering the way drugs act on the body are referred to as "altered drug action." Genetic conditions transmitted as single factors altering the way the body acts on drugs are referred to as "altered drug metabolism". These pharmacogenetic conditions can occur either as rare defects or as polymorphisms. For example, glucose-6-phosphate dehydrogenase deficiency (G6PD) is a common inherited enzymopathy in which the main clinical complication is haemolysis after ingestion of oxidant drugs (anti-

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malarials, sulfonamides, analgesics, nitrofurans) and consumption of fava beans.

As an illustrative embodiment, the activity of drug metabolizing enzymes is a major determinant of both
5 the intensity and duration of drug action. The discovery of genetic polymorphisms of drug metabolizing enzymes (e.g., N-acetyltransferase 2 (NAT 2) and cytochrome P450 enzymes CYP2D6 and CYP2C19) has provided an explanation
10 as to why some patients do not obtain the expected drug effects or show exaggerated drug response and serious toxicity after taking the standard and safe dose of a drug. These polymorphisms are expressed in two phenotypes in the population, the extensive metabolizer (EM) and poor metabolizer (PM). The prevalence of PM is
15 different among different populations. For example, the gene coding for CYP2D6 is highly polymorphic and several mutations have been identified in PM, which all lead to the absence of functional CYP2D6. Poor metabolizers of CYP2D6 and CYP2C19 quite frequently experience
20 exaggerated drug response and side effects when they receive standard doses. If a metabolite is the active therapeutic moiety, PM shows no therapeutic response, as demonstrated for the analgesic effect of codeine mediated by its CYP2D6-formed metabolite morphine. The other
25 extreme are the so called ultra-rapid metabolizers who do not respond to standard doses. Recently, the molecular basis of ultra-rapid metabolism has been identified to be due to CYP2D6 gene amplification.

Thus, the activity of Tango-77 protein, expression
30 of Tango-77 nucleic acid, or mutation content of Tango-77 genes in an individual can be determined to thereby select appropriate agent(s) for therapeutic or prophylactic treatment of the individual. In addition, pharmacogenetic studies can be used to apply genotyping
35 of polymorphic alleles encoding drug-metabolizing enzymes

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to the identification of an individual's drug responsiveness phenotype. This knowledge, when applied to dosing or drug selection, can avoid adverse reactions or therapeutic failure and thus enhance therapeutic or prophylactic efficiency when treating a subject with a Tango-77 modulator, such as a modulator identified by one of the exemplary screening assays described herein.

4. Monitoring of Effects During Clinical Trials

Monitoring the influence of agents (e.g., drugs, compounds) on the expression or activity of Tango-77 (e.g., the ability to modulate aberrant inflammation) can be applied not only in basic drug screening, but also in clinical trials. For example, the effectiveness of an agent, as determined by a screening assay as described herein, to increase Tango-77 gene expression, increase protein levels, or upregulate Tango-77 activity, can be monitored in clinical trials of subjects exhibiting decreased Tango-77 gene expression, decreased protein levels, or downregulated Tango-77 activity. Alternatively, the effectiveness of an agent, as determined by a screening assay, to decrease Tango-77 gene expression, decrease protein levels, or downregulate Tango-77 activity, can be monitored in clinical trials of subjects exhibiting increased Tango-77 gene expression, increased protein levels, or upregulated Tango-77 activity.

For example, and not by way of limitation, genes, including Tango-77, that are modulated in cells by treatment with an agent (e.g., compound, drug or small molecule) which modulates Tango-77 activity (e.g., as identified in a screening assay described herein) can be identified. Thus, to study the effect of agents on cellular proliferation disorders, for example, in a clinical trial, cells can be isolated and RNA prepared

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and analyzed for the levels of expression of Tango-77 and other genes implicated in the disorder. The levels of gene expression (i.e., a gene expression pattern) can be quantified by Northern blot analysis or RT-PCR, as described herein, or alternatively by measuring the amount of protein produced, by one of the methods as described herein, or by measuring the levels of activity of Tango-77 or other genes. In this way, the gene expression pattern can serve as a marker, indicative of the physiological response of the cells to the agent. Accordingly, this response state may be determined before, and at various points during, treatment of the individual with the agent.

In a preferred embodiment, the present invention provides a method for monitoring the effectiveness of treatment of a subject with an agent (e.g., an agonist, antagonist, peptidomimetic, protein, peptide, nucleic acid, small molecule, or other drug candidate identified by the screening assays described herein) comprising the steps of (i) obtaining a pre-administration sample from a subject prior to administration of the agent; (ii) detecting the level of expression of a Tango-77 protein, mRNA, or genomic DNA in the preadministration sample; (iii) obtaining one or more post-administration samples from the subject; (iv) detecting the level of expression or activity of the Tango-77 protein, mRNA, or genomic DNA in the post-administration samples; (v) comparing the level of expression or activity of the Tango-77 protein, mRNA, or genomic DNA in the pre-administration sample with the Tango-77 protein, mRNA, or genomic DNA in the post administration sample or samples; and (vi) altering the administration of the agent to the subject accordingly. For example, increased administration of the agent may be desirable to increase the expression or activity of Tango-77 to higher levels than detected,

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i.e., to increase the effectiveness of the agent.
Alternatively, decreased administration of the agent may
be desirable to decrease expression or activity of
Tango-77 to lower levels than detected, i.e., to decrease
5 the effectiveness of the agent.

C. Methods of Treatment

The present invention provides for both
prophylactic and therapeutic methods of treating a
subject at risk of (or susceptible to) developing or
10 having a disorder associated with aberrant Tango-77
expression or activity. Alternatively, disorders
associated with aberrant IL-1 production can be treated
with Tango-77. Such disorders include acute and chronic
inflammation, asthma, some classes of arthritis,
15 autoimmune diabetes, systemic lupus erythematosus and
inflammatory bowel disease.

1. Prophylactic Methods

In one aspect, the invention provides a method for
preventing in a subject, a disease or condition
20 associated with an aberrant Tango-77 expression or
activity (or aberrant IL-1 expression or activity), by
administering to the subject an agent which modulates
Tango-77 expression or at least one Tango-77 activity.
Subjects at risk for a disease which is caused or
25 contributed to by aberrant Tango-77 expression or
activity can be identified by, for example, any or a
combination of diagnostic or prognostic assays as
described herein. Administration of a prophylactic agent
can occur prior to the manifestation of symptoms
30 characteristic of the Tango-77 aberrancy, such that a
disease or disorder is prevented or, alternatively,
delayed in its progression. Depending on the type of
Tango-77 aberrancy, for example, a Tango-77 agonist or
Tango-77 antagonist agent can be used for treating the

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subject. The appropriate agent can be determined based on screening assays described herein.

2. Therapeutic Methods

Another aspect of the invention pertains to methods of modulating Tango-77 expression or activity for therapeutic purposes. The modulatory method of the invention involves contacting a cell with an agent that modulates one or more of the activities of Tango-77 protein activity associated with the cell. An agent that modulates Tango-77 protein activity can be an agent as described herein, such as a nucleic acid or a protein, a naturally-occurring cognate ligand of a Tango-77 protein, a peptide, a Tango-77 peptidomimetic, or other small molecule. In one embodiment, the agent stimulates one or more of the biological activities of Tango-77 protein. Examples of such stimulatory agents include active Tango-77 protein and a nucleic acid molecule encoding Tango-77 that has been introduced into the cell. In another embodiment, the agent inhibits one or more of the biological activities of Tango-77 protein. Examples of such inhibitory agents include antisense Tango-77 nucleic acid molecules and anti-Tango-77 antibodies. These modulatory methods can be performed *in vitro* (e.g., by culturing the cell with the agent) or, alternatively, *in vivo* (e.g., by administering the agent to a subject). As such, the present invention provides methods of treating an individual afflicted with a disease or disorder characterized by aberrant expression or activity of a Tango-77 protein or nucleic acid molecule. In one embodiment, the method involves administering an agent (e.g., an agent identified by a screening assay described herein), or combination of agents that modulates (e.g., upregulates or downregulates) Tango-77 expression or activity. In another embodiment, the method involves

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administering a Tango-77 protein or nucleic acid molecule as therapy to compensate for reduced or aberrant Tango-77 expression or activity.

Stimulation of Tango-77 activity is desirable in situations in which Tango-77 is abnormally downregulated and/or in which increased Tango-77 activity is likely to have a beneficial effect. Conversely, inhibition of Tango-77 activity is desirable in situations in which Tango-77 is abnormally upregulated and/or in which decreased Tango-77 activity is likely to have a beneficial effect.

This invention is further illustrated by the following examples which should not be construed as limiting. The contents of all references, patents and published patent applications cited throughout this application are hereby incorporated by reference.

EXAMPLES

Example 1: Isolation and Characterization of Human Tango-77 cDNAs

Cytokine genes IL-1 α , IL-1 β and IL-1ra have been found to be closely clustered on chromosome 2, i.e., IL-1 α , IL-1 β and IL-1ra are located within 450 kb of each other. BAC clones containing IL-1 α and IL-1 β were used to identify other proximal unknown cytokine genes. To do this, a BAC clone containing IL-1 α and IL-1 β was selected from a BAC library (Research Genetics, Huntsville, Alabama) using specific primers designed against IL-1 α and IL-1 β . The DNA from the BAC was extracted and used to make a random-sheared genomic library. From this BAC library, 4000 clones were selected for sequencing. The resulting genomic sequences were then assembled into contigs and used to screen proprietary and public data bases. One genomic contig was found to contain two

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segments of sequences which resemble IL-1ra. These two segments are potential exons of Tango-77 gene.

Two PCR primers were then designed from the two potential exons and used to screen a panel of cDNA libraries for the expression of a Tango-77 message. A cDNA library from TNF- α treated human lung epithelia showed a positive band of the predicted size (i.e., if the two exons are spliced together). Using the PCR fragment as a probe, a single cDNA clone was isolated from the same library. This cDNA contains an insert of 989 bp. The cDNA clone contains three possible open reading frames. The first open reading frame encompasses 534 nucleotides (nucleotides 356-889 of SEQ ID NO:1; SEQ ID NO:3) and encodes a 178 amino acid protein (SEQ ID NO:2). This protein may include a predicted signal sequence of about 63 amino acids (from amino acid 1 to about amino acid 63 of SEQ ID NO:2 (SEQ ID NO:4)) and a predicted mature protein of about 115 amino acids (from about amino acid 64 to amino acid 178 of SEQ ID NO:2 (SEQ ID NO:5)).

The second putative nucleotide open reading frame encompasses 498 nucleotides (nucleotides 389-889 of SEQ ID NO:1; SEQ ID NO:6) and encodes a 167 amino acid protein (SEQ ID NO:7). This protein includes a predicted signal sequence of about 52 amino acids (from amino acid 1 to about amino acid 52 of SEQ ID NO:7 (SEQ ID NO:8)) and a predicted mature protein of about 115 amino acids (from about amino acid 53 to amino acid 167 of SEQ ID NO:7 (SEQ ID NO:9)).

The third open reading frame (nucleotides 372-889 of SEQ ID NO:1; SEQ ID NO:10) encompasses 408 nucleotides and encodes a 136 amino acid protein (SEQ ID NO:11). This protein includes a predicted signal sequence of about 21 amino acids (from amino acid 1 to about amino acid 21 of SEQ ID NO:11 (SEQ ID NO:12)) and a predicted

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mature protein of about 115 amino acids (from about amino acid 22 to amino acid 136 of SEQ ID NO:11 (SEQ ID NO:13)).

Tango-77 is predicted to be 35% identical to human IL-1ra at the amino acid level.

Example 2: Expression of Tango-77 mRNA in Human Tissues

The expression of Tango-77 was analyzed using Northern blot hybridization. A PCR generated 989 bp Tango-77 product was radioactively labeled with ³²P-dCTP using the Prime-It kit (Stratagene; La Jolla, CA) according to the instructions of the supplier. Filters containing human mRNA (MTNI and MTNII: Clontech; Palo Alto, CA) were probed in ExpressHyb hybridization solution (Clontech) and washed at high stringency according to manufacturer's recommendations.

Tango-77 mRNA was not detected in any unstimulated tissues (brain, liver, spleen, skeletal muscle, testis, pancreas, heart, kidney and peripheral blood leukocytes) mRNA on Clontech Northern blots.

Over 96 cDNA libraries were then tested for the presence of Tango-77 using PCR amplification. Only three libraries displayed a positive signal. These libraries were the TNF α -treated bronchoepithelium, TNF α -treated SSC cell line and anti-CD3-treated T cells.

Example 3: Characterization of Tango-77 Proteins

In this example, the predicted amino acid sequence of human Tango-77 protein was compared to the amino acid sequence of known protein IL-1ra. In addition, the molecular weight of the human Tango-77 proteins was predicted.

The human Tango-77 cDNA (Figure 1; SEQ ID NO:1) isolated as described above encodes a 178 amino acid protein (Figure 1; SEQ ID NO:2) or a 167 amino acid

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protein (Figure 1; SEQ ID NO:7) or a 136 amino acid protein (Figure 1; SEQ ID NO:11). The signal peptide prediction program SIGNALP Optimized Tool (Nielsen et al. (1997) *Protein Engineering* 10:1-6) predicted that

5 Tango-77 includes a 63 amino acid signal peptide (amino acid 1 to about amino acid 63 of SEQ ID NO:2 (SEQ ID NO:4)) preceding the 115 mature protein; or preceding the 115 mature protein (about amino acid 52 to amino acid 167 of SEQ ID NO:7 (SEQ ID NO:8)); or preceding the 115

10 mature protein (about amino acid 21 to amino acid 136 of SEQ ID NO:11;SEQ ID NO:12).

As shown in Figure 2, Tango-77 has a region of homology to IL-1ra (SEQ ID NO:14).

Mature Tango-77 has a predicted MW of about 13 kDa

15 and the predicted MW for the immature Tango-77 is 19.6 kDa, 18.5 kDa or 15.2 kDa, not including post-translational modifications.

Example 4: Preparation of Tango-77 Proteins

Recombinant Tango-77 can be produced in a variety

20 of expression systems. For example, the mature Tango-77 peptide can be expressed as a recombinant glutathione-S-transferase (GST) fusion protein in *E. coli* and the fusion protein can be isolated and characterized. Specifically, as described above, Tango-77 can be fused

25 to GST and this fusion protein can be expressed in *E. coli* strain PEB199. Expression of the GST-Tango-77 fusion protein in PEB199 can be induced with IPTG. The recombinant fusion protein can be purified from crude bacterial lysates of the induced PEB199 strain by

30 affinity chromatography on glutathione beads.

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Example 5: Alternatively spliced forms of IL-1ra and
Tango-77

Computer program Procrustes (Gelfand et al., 1996, *Proc. Natl. Acad. Sci. USA*, 93:9061-9066) is an alignment
5 algorithm that predicts the presence of alternatively
spliced exons for a protein of interest in a stretch of
genomic DNA. Using the IL-1ra sequence, Procrustes was
used to search for the presence of additional sequences
that might encode for alternatively spliced forms of IL-
10 1ra in the two overlapping BAC genomic sequences (see
Fig. 3 and Fig. 4). Potential sequences that encode
variant exons for IL-1ra were identified. These
predicted exons aligned well with the N-terminal region
of IL-1ra, but were not present in Tango-77. The results
15 from Procrustes predicts the existence of more spliced
forms of IL-1ra.

Furthermore, Procrustes also predicted an
additional sequence in BAC1 and BAC2 that encodes an
alternatively spliced exon for Tango-77 (T77-procrustes;
20 Fig. 5). This predicted splice variant form of Tango-77,
T77-procrustes, was aligned with Tango-77 (Fig. 6) and
with IL-1ra and IL-1 β (Fig.7).

PCR primers within this sequence can be used to
generate a product that can be used to screen a panel of
25 cDNA libraries using standard techniques. Suitable cDNA
libraries include libraries made from TNF α -treated
bronchoepithelium, TNF α -treated SSC cell line and anti-
CD3-treated T cells. The resulting cDNA clone(s) can be
isolated from the library and sequenced to identify
30 additional Tango-77 cDNAs.

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Equivalents

Those skilled in the art will recognize, or be able to ascertain using no more than routine experimentation, many equivalents to the specific
5 embodiments of the invention described herein. Such equivalents are intended to be encompassed by the following claims.

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What is claimed is:

1. An isolated nucleic acid molecule selected from the group consisting of:

- a) a nucleic acid molecule comprising a
5 nucleotide sequence which is at least 45% identical to the nucleotide sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807, or a complement thereof;
- 10 b) a nucleic acid molecule comprising a fragment of at least 300 nucleotides of the nucleotide sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807, or a complement thereof;
- 15 c) nucleic acid molecule which encodes a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the
20 plasmid deposited with ATCC as Accession Number 98807;
- d) a nucleic acid molecule which encodes a fragment of a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID
25 NO:12, SEQ ID NO:13, wherein the fragment comprises at least 15 contiguous amino acids of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or the polypeptide encoded by the cDNA insert of the plasmid
30 deposited with ATCC as Accession Number 98807; and
- e) a nucleic acid molecule which encodes a naturally occurring allelic variant of a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9,

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SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807, wherein the nucleic acid molecule hybridizes to a nucleic acid molecule comprising SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10, or the complement thereof under stringent conditions.

2. The isolated nucleic acid molecule of claim 1, which is selected from the group consisting of:

10 a) a nucleic acid comprising the nucleotide sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, or SEQ ID NO:10 or the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807, or a complement thereof; and

15 b) a nucleic acid molecule which encodes a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807.

3. The nucleic acid molecule of claim 1 further comprising vector nucleic acid sequences.

4. The nucleic acid molecule of claim 1 further comprising nucleic acid sequences encoding a heterologous polypeptide.

5. A host cell containing the nucleic acid molecule of claim 1.

6. The host cell of claim 5 which is a mammalian host cell.

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7. A non-human mammalian host cell containing the nucleic acid molecule of claim 1.

8. An isolated polypeptide selected from the group consisting of:

- 5 a) a fragment of a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, wherein the fragment comprises at least 15 contiguous amino acids of SEQ ID
10 NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, or SEQ ID NO:13.
- b) a naturally occurring allelic variant of a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8,
15 SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807, wherein the polypeptide is encoded by a nucleic acid molecule which hybridizes to a nucleic acid molecule
20 comprising SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, SEQ ID NO:10 or the complement thereof under stringent conditions;
- c) a polypeptide which is encoded by a nucleic acid molecule comprising a nucleotide sequence which is
25 at least 55% identical to a nucleic acid comprising the nucleotide sequence of SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, or SEQ ID NO:10.

9. The isolated polypeptide of claim 8 comprising the amino acid sequence of SEQ ID NO:2, SEQ ID
30 NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807.

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10. The polypeptide of claim 8 further comprising heterologous amino acid sequences.

11. An antibody which selectively binds to a polypeptide of claim 8.

5 12. A method for producing a polypeptide selected from the group consisting of:

a) a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID
10 NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807;

b) a fragment of a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID
15 NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807, wherein the fragment comprises at least 15 contiguous amino acids
20 of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the plasmid deposited with ATCC as Accession Number 98807; and

25 c) a naturally occurring allelic variant of a polypeptide comprising the amino acid sequence of SEQ ID NO:2, SEQ ID NO:4, SEQ ID NO:5, SEQ ID NO:7, SEQ ID NO:8, SEQ ID NO:9, SEQ ID NO:11, SEQ ID NO:12, SEQ ID NO:13, or an amino acid sequence encoded by the cDNA insert of the
30 plasmid deposited with ATCC as Accession Number 98807, wherein the polypeptide is encoded by a nucleic acid molecule which hybridizes to a nucleic acid sequence of

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SEQ ID NO:1, SEQ ID NO:3, SEQ ID NO:6, or SEQ ID NO:10
under stringent conditions;

comprising culturing the host cell of claim 5
under conditions in which the nucleic acid molecule is
5 expressed.

13. A method for detecting the presence of a
polypeptide of claim 8 in a sample, comprising:

- a) contacting the sample with a compound which
selectively binds to a polypeptide of claim 8; and
- 10 b) determining whether the compound binds to the
polypeptide in the sample.

14. The method of claim 13, wherein the compound
which binds to the polypeptide is an antibody.

15 15. A kit comprising a compound which selectively
binds to a polypeptide of claim 8 and instructions for
use.

16. A method for detecting the presence of a
nucleic acid molecule of claim 1 in a sample, comprising
the steps of:

- 20 a) contacting the sample with a nucleic acid
probe or primer which selectively hybridizes to the
nucleic acid molecule; and
- b) determining whether the nucleic acid probe or
primer binds to a nucleic acid molecule in the sample.

25 17. The method of claim 16, wherein the sample
comprises mRNA molecules and is contacted with a nucleic
acid probe.

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18. A kit comprising a compound which selectively hybridizes to a nucleic acid molecule of claim 1 and instructions for use.

19. A method for identifying a compound which
5 binds to a polypeptide of claim 8 comprising the steps of:

- a) contacting a polypeptide, or a cell expressing a polypeptide of claim 8 with a test compound; and
- 10 b) determining whether the polypeptide binds to the test compound.

20. The method of claim 19, wherein the binding of the test compound to the polypeptide is detected by a method selected from the group consisting of:

- 15 a) detection of binding by direct detecting of test compound/polypeptide binding;
- b) detection of binding using a competition binding assay; and
- c) detection of binding using an assay for
20 Tango-77-mediated signal transduction.s

21. A method for modulating the activity of a polypeptide of claim 8 comprising contacting a polypeptide or a cell expressing a polypeptide of claim 8 with a compound which binds to the polypeptide in a
25 sufficient concentration to modulate the activity of the polypeptide.

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22. A method for identifying a compound which modulates the activity of a polypeptide of claim 8, comprising:

- a) contacting a polypeptide of claim 8 with a
5 test compound; and
- b) determining the effect of the test compound on the activity of the polypeptide to thereby identify a compound which modulates the activity of the polypeptide.

GTGGACCCACGGCTCCGCAGACGTCTACCTGGGGGTCCCGTCTGCGCTCCCGGGATGGAACGCCAGGGGAACTTA 79
GGCAGGGGAGCGGACGGGCACCTCCCGCGGGACGAACTCACTCGGTGGCCCTCCTACTTCCCGGGCCGTGTTCCAACGCC 158
TGAGAATAACGGGAACAGCGGTCTACTCACCAGACAGCGGCAGCAGCGGCCTCTCTCAATTGGGCAAAGCACTCCAGAC 237
TCTTTTGGAAAGAGTGACACCAAAGGCAAGCACCTGCTTGGCAGGCCCCCTCAGCTTCTACGCAAGTATAAGTCTTGGACTT 316
CATTCCATTTTCTGTTGAGTAATAAACTCAACGTTGAAA M S F V G E N S G V 10
ATG TCC TTT GTG GGG GAG AAC TCA GGA GTG 385
K M G S E D N E K D E P Q C C L E D P A 30
AAA ATG GGC TGT GAG GAC TGG GAA AAA GAT GAA CCC CAG TGC TGC TTA GAA GAC CCG GCT 445
G S P L E P G P S L P T M N F V H T K I 50
GGA AGC CCC CTG GAA CCA GGC CCA AGC CTC CCC ACC ATG AAT TTT GTT CAC ACA AAG ATC 505
F F A L A S S L S S A S A E K G S P I L 70
TTC TTT GCA TTA GCC TCA TCC TTG AGC TCA GCC TCT GCG GAG AAA GGA AGT CCG ATT CTC 565
L G V S K G E F C L Y C D K D K G Q S H 90
CTG GGG GTC TGT AAA GGG GAG TTT TGT CTC TAC TGT GAC AAG GAT AAA GGA CAA AGT CAT 625
P S L Q L K K E K L M K L A A Q K E S A 110
TCA TCC CTT CAG CTG AAG AAG GAG AAA CTG ATG AAG CTG GCT GCC CAA AAG GAA TCA GCA 685
R R P F I F Y R A Q Y G S W N M L E S A 130
CGC CCG CCC TTC ATC TTT TAT AGG GCT CAG GTG GGC TCC TGG AAC ATG CTG GAG TCG GCG 745
A H P G W F I C T S C N C N E P V G V T 150
GCT CAC CCC GGA TGG TTC ATC TGC ACC TCC TGC AAT TGT AAT GAG CCT GTT GGG GTG ACA 805
D K F E N R K H I E F S F Q P V C K A E 170
GAT AAA TTT GAG AAC AGG AAA CAC ATT GAA TTT TCA TTT CAA CCA GTT TGC AAA GCT GAA 865
M S P S E V S D * 179
ATG AGC CCC AGT GAG GTC AGC GAT TAG 892
GAAACTGCCCCATTGAACGCCCTTCTCGCTAATTTGAACTAATTGTATAAAAAACCCAAACCTGCTCACTAAAAA 971
AAAAAAAGGGCGGCCGC 989

Fig. 1

1
 IL1ra-human 50
 MEICRGLRSH LITLLFLFH SETICRPSGR KSSKMQAFRI WDVNQKTFYL
 T77-human
 IL1b-human
 Consensus
 51
 IL1ra-human 100
 RNNQLVAGYL QGPNVNLEEK IDVVFIEPH. ALFLGIHGKK MCLSCVKSGD
 T77-human
 IL1b-human
 Consensus
 101
 IL1ra-human 150
 ETR..LQLEA VNITDLSENR KQDKR.FAFI RSDSGPTTSF ESAACPGWFL
 T77-human
 IL1b-human
 Consensus
 151
 IL1ra-human 192
 CTAMEADQPV SLTNMPDEGV MVTKFYFQED E-
 T77-human
 IL1b-human
 Consensus

FIG. 2

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>Contig1
GAAGTGAAGATATAATGTATAGTAGTAATATATAATGTTAGGTGAATTAA
AGGAAATAGAATATATTGGGGAGTAATTATGGGTGTAAAGAAATATAGTA
GGGAAGTATTTAGATTTGAGAAAAAAGGAATTTAGTGTAGGTGAA
NAATAAAAGNANAAGGTTAAAAAATTAATAAATAAATAAATAAATAA
AAATAAAATAAATAAATAAATAAATAAATAAATAAATAAATAAATAA
AAAAATAAGAAATGGAAGTGGATTCTTAGAAAAAAGAAAGTAAGGTGA
TATGAGGAGATAGAGAGGATGTGGTGTGAGATGATTGGTTAATTAGAAA
ATAGGTTTTGAATAGAGTGGGAAAGTAGAGTTTGGTAAATGTGGGGGA
AGAGGGTAATGTTGTTTGAAGTGAAGAAAAAATGGTATATTTTATAAAA
TAATGAGGAAAGTGTGTGAAAAAATTTGGGGTGGGGTGGGGTGGGGTGG
ATATAAAGTTGTGGAAAAATTTGGGGGTGGGGTGGGGTGGGGTGGGGTGG
GTTATTTAAAGAATGAAATGAATTTTGTGTTGTAATTTGGGGATAAGAA
ATTAATGTTTAGAAAGAAAGGGAAGAAATTTGAAGAAAAAATTTAGATTT
TGGAAATTTAAAAATATTGTGGGTGTAAATAGGAAGGATTTTAAAGGTA
ATTGTGGAAGGGATTTGTGTGGAAAAATAATAGGGAGAAAAATGGGG

>Contig2
GCATCTAACTGGAGCCTGCATTATTACAGATTTAGCATCACCAAAGTCTA
AACAAATTAGACTGACTAAGGCAGAACTGCCCTTATGACAGCAGACATAAG
AAGGAAAAGGCCAAAACACTGTGTTAAAAATTATCCAAATGTGAGGAAAA
GGCAAAGAGAGTAGGTGTGCCTTTTGTGTTCTAAGCTGCCTGCCCAAGG
GGCATCTGATGCTCTCAGGCAGGAGTCCACAAATTTTGTAAAAGA
TCAGATAGTAAATCTTTTCAGCGTGAAGAGCATGAGGTCTCTGTACAAA
TACTCAACCACCATTACAACATGAAAGCAGCCAACAGACAACACATGACA
AATGAGTGTGGCTGTGTTCCAGTAAATCTTGATTACAAAAACAGGCAAGA
GGCCAGAGCTGACCCATGGGCCATAGTTTGTGACCCCTTCTGTAAAGGA
AAGTATTTTGTGTTGACTGTGTTTACCATTGATTGAACACAAGGCTCT
GTAAAGTTACTTGTAACTTGCAGAAGATTGATGAGTGGCAAGTAATTTT
TATTCACCAGAATATAAAATTATTTCTGTTTCAAGTAGAAAAGATAAACCAA
CTGTGATATTATGGTCTG

>Contig3
GGGGTGTCTGTCTACCATGTGCTCGCAGTTCTGTAATAAATGTTCTCTCA
AGATCCTTAAAAATCTCTTGGAAATTATAAAAAATATTGGAAAGAGAAGAAC
AGTTTAAATATATATATATATATATATATTTTTTTTGTAGATGGAGTCTT
GCTCTGTGCTCCAGGCTGGAGTGCAGTGGCGCAAACCTTGGTTCCACCACAA
CCTCTGCCTCCCGGTTCAAGCGATTCTTCTGCCTCAGCCTCCTGAGTAG
CTGGGACTACAGGCGCCCGCCACCACGCCCAGCTAATTTTGTATTTTAA
GTAGAGACGAGGTTTTACTATGTTGGCTAGGCTGGTCTCAAACCTCCTGAC
CTTGTGATCTGCCCGCTTGGCCTCCCAAAGTGTGGGATTACAGGTGTG
AGCCACTGCACCTGGCCAGTTTTTAAATATATTTTTTAAAAACACTTGAA
TAAGAGTCAAGTGTAACTAGAAGTTTAAAAATGCTTCACAGAACACCCAG
GGTTTACATTACAAGATTCTCACAACAAACCTATTGTAAAGGTGAGTAAG
GCATGTTATTACAGAGAAAAAGTTTGGGAGCAAACTGTAAAAAATTATAT
TTTTGTTGATTTTCTAAGAGAAAGAGTATTGTTATGTTCTCCTAACCTC
TGTTGATTACTACTTTAAGTGATTTCTTGGAGGCACATGATGATCC

>Contig4
GCCGTTTCATAGAAAACTGAAAGCAATAAGATGACTAGGTAAGCATGACAT
TTAAAAGGTATTCATGGGACGTGGTTACAAAACCAACTCACAACATAAAA
GTCTTAGGACCTCTCGCTGACTTAGGAGCCTGATCCCAACTCTGAGAATG
ACTCAGTGTGTTACCCTGTGGCTAGTGTAGACCAATGATCCTGTCTCAGA
GTCAC TAGCCAACAGCCCATATCAAGTACTTGAAACTTTGACTCAGAAAC
CTCAGTGTGAGAACCTTTGACCTAGGAACCACTGTAGTGGTTAACTGCA
ATTTGCACCCCTTAGTTGAGGCTTTACAACACCGGGGGCGGGGAGGGGA
AAGGCATANANCTGATGACCTAAAGGAAACCCATTGCAGCAACGCTTTTG
TGTTAAGTGTAATAAAGTGTGTTTGTAGAACTCTCCAGGTAATGCCTT
TGTTATTTAATGTGTCTGAGACAATTCTGCACATTAAAGAAATAAAAAA
TTACCTTTGTAATTTCAATTTGAAATGTGTAATTGACATTAGACTTCTATT
TGAATTTGAAATGTCTAAACAATGTGGTTAAGTTTGTAAAAGGTGTGTG
AATTTTGAAGTCTGATTACTACATTTTTTTTAAATTTCTTTTTTTTTGG
AGTTTTAGGGATTGCTTAGATGGCTAGAAAGATTTTATTCATCAGATTTT

FIG. 3 (1 of 52)

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TAAGTCTGCCCTTGGCAGGCACCTTGACAG. JTTTGAAAGAATCAGATATATC
AAATTTGTAGTTTAAAAATATTTAAGGGAACCTCAATTAACCTATGCTAGAAA
AGAGAATTAAGTATTTAGGAGGATTTAATATGGTGTGAAAGTTGTGAAAA
TCAAATGGAGACACTAATGTTAAGAAAACCTGATAAATGGAACCAGGG
AAAGGCATGAAGATAGAGTTCTCACACTTGTATCCCTGATCATGAAAAAG
ATCTGC

>Contig5

GGGTTTTTCCGCGTTTTTACCCGAAATCTTCAAGGGATGGGAAAAAGAAA
ATTGCTAAAAAATCTCGGTTTTTTGGTTTTTAACAGATATTTACACNTGG
ATCCCATTTATTATGTTGTCCCCAAGGTTTTTCGGTGGGTTCCCAATCAGT
TAGCCCCCTCCACAGTGAAAGCACTTTACTTTATCACCTTCACCTAAAG
CATAAAATCCAGCTCTTGAAAGCTGCTCCTTGTTAACTGAATATATCCAC
ATCCCAAAAGTAATGATCCATGCTTCATAATCTGCCACGGATGGATGGAT
GGATGGATGGATGGATGGATGGATGGATGAATGGATGGATTGATTTCTTG
GAGGATTTGTTGAATTTGGGAAATTCACGCCAGGACAGCTGGCCCAAAC
TGCCCGGACAATCTGCTCGGTACAAGGGGAGGGTCTTGAGAGGGTGGC
GCCCGAGCCCCAGTTTGGAAATGCCAATCTGGCTCTGCAGCCGGGCCTTA
GCCACTTGGGTCTGGCGTCCCTCCATTATTAGCGCCATGCCGGCTCGGG
TGCTGCCAAGTCCCTGAGAGCACAAAGCC

>Contig6

CGCGCTCAAGAAAAGCTGAAGTGTGAATGTTCTGTCTACCTTCACAGTAA
ATGCTAAGAGAATGACCCAGAGCAGAGGGTATCACTCTGCTACGGAGGA
TTGATTGTAAGTGGCTCTCCTGCCCTTAGCAAGAAATGCCAGAACCATGGT
CATTCAAGTCTTGACCAAAACTGCCTTCATGAGAATCAACTTCCCCAA
GAAAAAAAAGCAGAAACAGGCAAGCTTCCAGCATGGTAGGTAATACTG
ACCTTCTTCCCTCCTTCTTGGAGATTACACAGTAATAATGCATAAA
GCTTTGCCAATGGACTAAGCACTGCCAGGGGTTTTTGTCTGCCTGGAC
TGAAATGCTCTTTTTGCGTTATCATAGAATCCAGTGCAGTCTGAGTAGA
CTCTAAGCAAAAGGGACATTTTTCAAAAAGGCTTTAAATTGCTAGTACAA
AGAAGGCAACAAAACCTTGGCTAAGTGTGGACAGATTAACCTCACTTGGTGT
TTTGGCTCTTCAGTTTTCCCTTGGCTGCCAAGTACTCCTGAAGCTTTCTC
TGCGGCTCTTCTGCAAGCAGGCAAGCAAAAAACGACTGAACCTTTATTT
CGAGAT

>Contig7

GAAGAGCCGCTAACTTGCTGTAGTGATAAGGAATGAACTAAGGCTAGGGA
CATATTAACATCCGCTGGTGGTGAAGTCTTTAGCCTAGATCTTACCCCACT
CCTGCTCCTTCCATATGGTTCGGTCTCAGGCTCACTACCGATCAATGGCG
TACTAAAAGCACTAACTATAGACTCCAACACGTCTGTCTGTGTTTACAG
ACAAGCCCTGGAGTTAATCCCTCTGACAGTAGCTCAGATAAGGATGGGCT
ATCATGGGCCCCGGAACCTGGGGCATGACGCTCGTCACCAACGCATGAGCTC
CCCAAGTATGCTATACCTGTCCCTATGAAGGGCTTCCAACCTCTATGTGCA
GTCCCCATGTGGAGAGTCAGGTATTGATTGATCAAGCCAGGGGTGTGGTG
AATGGGGAGCTTCTACAGGGGTAATGATAATTGAAATGCACGGTGATGG
GGATTTTCATATTGGTCTCCTAAGGAGATAACAGATTGGATGCGGGGTGCG
ATATTCCACTGCCAGGGTGTGTACCGAGGGTATCTGCAGGTGGATCTCC
TCCCCACGTTTGATTAACTCCTGTCTTGGGAAGCATAGACGGGCGGGG
GAAATGATGAAGGTGACCACTCCCC

>Contig8

GGGAACGCAGTGCTCTGTACGATGGCCTTGATTGCGAATTCCTGCAGGGG
GGG

>Contig9

GGCAAGAGATTTAATATTCATTCCATCTTCATTTGGAAGATGAAAAATTG
GGGACCAGAGAGGGGAGGGGACTGGGCCAAGTTTTCAAGAAAAGTCAGT
AGGAATTGTGAATTCCTGGGGGCCGGGGCCATTAGTGCTGTTTTGGATC
AGTAAATGGAGATGTGAGTTTCAACAGTAACAGGGACATTTTAAATTA
AATGATTAACTTTAGAAAATGTCCTATTTTGTAAATAATGATGGATTCA
CAGGAAGGTACAAAGAAATGTCCAGAGAGTTCNTGAGCCCCCTTCAGCCA
GCTTCTTCCAATGTTAACATCTTGCAATTATTATAGTACAACATCAAACT
GGGAAATC3ATATTGGTACTGTCCAGATAGCTTACTCAGATTTTGCCAGT
TATACTTCCACTCATTTGTGTGTGTGTGTGTGTGTGTGTGTGTGTGTGTG

FIG. 3 (2 of 52)

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TGTGTGTAGCTCTATGCAATTTTATG1...GTAGCTTCATGTAACCAAC
AATCACAATACTTAACCTATGCCCTCATCAAGACTCTCTCTTGCTATGC
TTTACAGCTGTATCCTCTTCATCTCCAAACCCTAAGCCCACCTCACCGCC
TCCACCATCTCTAATCCCTGGCAACCACTATTCTGTGCTCCATCTCTGTA
ATTAATTGTGTTAATTAATGTTATACAAATGGAATCATGAAGTATGTGTC
CTTTGAGATTGGGCTGTTAATTTTCACTCAGCACAATTTCCGTGAGTCT
AATCCAACCTGTGTGTAGCAGTAATTTCTTCTTATTATTGCTGAATAAT
ATGCCATGGTATGGATGTATCACAGTGTGTCTAATCCTTTGCCATTGAA
AGGAATTTGGATAATTTCCAGGTTTTGGCTATTATGAATAAAGTGAACAT
AAGACATGTGTGTACAAATTTTGGTGTGATCAAAAGTCTCATTCTCTGG
GATAAATGCCCGGTAATGAAATGGCTGGGTTGTGTGGG

>Contig10

GCAAGAACACAGGCGCGTATTATAACCTTACTACCAAGACCTGAACCCAT
ATAAAGGTTTATGCGTAACAATCATCATCCCTGTTCCAGAAGATTACACG
TACGACCACGCTGGCTCACCAGCTCACGTGGGCCAGTACCAGAAATTCT
CCCAAACAAACAGTCGTGTCTGAAAACAATCGCGGTGACCTCCACGGTTA
GAAAAGCCTGTTTTCAAGTCTTGAATTGCCACATATTAGCTGGGTAAT
TTGGGCATCACATTTACTCTCTCCGAATTTGAGATTGCAAAAACCTCATT
GATTGTTTTGTGGATTGAAAGAAATAATGTAAATTTAGGCCGAGTGCTTT
GACTTACGCTGTAAATCCTATCACTTTGGGAGGCCAAAGCAGGAGGGTCA
CTTGAGCTCAGGAATTTGAGACCACCTCTGGCAACATAGTGAGATCCTGT
CTCTACAAAAAATTTTTTTTAAATTATCCAGCATGGTGGTACACGCCTGT
ATTTCCAGCTACTCAGGAGACTGAGGTGTGAGGATTGCTAGAACCCTGGGA
GATCAAGTCAACAGTGAGCCGTGGTTGTGCCACTGCCCTCCAACCTCAGT
GACAGAGGAAGACCCTGTCTCAAAAAAAAAAAAAAAAAAGTAGTAAGTTTAA
AGAACTTAGTGTAGGCCTGGCATATAAATGATATTGTTGATGTTGATGTT
AGCTTGAAGGCACATTTATAGGAGTAGGGATTTTATAACATTATGAGCCT
GAGAGCACATATAATGTTCCC

>Contig11

GGTCTAACATGCTCCAACTGAAGAAACCCACACTTGTCCGGCAAGGAAA
CTACTACAGATTTCTGACCTACTGTGCAATTCGGGGCATGCGACGGGAC
TGTGTTTTCTGGGTACGCTGTCTCAGGTTCTGTCTGGGATGTAAGAATTCAA
CTTCAGTAGTTCTCTCATAGACGCCGACGAGAGGGGCGTCTCTTTCTCT
GATGAATCTGCCAGATCTTCCACTTCATAGAGTCTAAATCCTCCGATTCTG
ATCTACTGGAGACCCCCAGTTACAAAAACGTCTAACGTCCGTGACAGCT
CCCCACATAGGGAAGATCACCTGAGTCTCACTACCTCACATTAGTGCTA
TCTCCAGCCCCATGCTATCTACGAGATGGTCACGCGAGGTTTAAGGGGTC
TCCGATTCCGGTGGTCCGATTGAGCTAATCGTGGCCCTACGTGAACGATC
ACTCCTGCTCGTAACATCGATACAGGGTCCGCGTGACAAATGGTACTACG
TAGGTTCTCAGGTCAATGCCGCGTCACGAATGAGCCTAACTACCCCATAA
GTGCACGTACTGTGTTACCTTCTCTGTTCCGGCAAACCTGCTACTGTATG
CTGTGCTTGTTT

>Contig12

AGGCTCCATGTGCTCTAGCCTGATTATCTTTTTCAAGTGTTTTATTGCTA
ATCTATAAGGCCCTTTTCGTAAAATGTTCACTCATTTTTCTAATTAGATAT
TTTTTTTAAATGTTGAGTTTGGAGAGTTCTTTAGATATTTTAGATACAAGT
CCATTGTCAAATATGTGATTACAAATATTTCTCTCAATCTGTAATTTA
GTTTTATCCTCTTAACAGGGTCTTTTGGAGAGCAAATAATTTGATTTTC
ATAAGGTTCAAATATTAATTTTTCTTGTATAGTTCACACTTCTAGTGT
TAAGTCTAAAAACTGTGCCTTGTCTATAGGTACCAAAGGTTTTCTCCAGTT
TTTTTTCTAGAAGTTTAGAGTTTTCATGTTTTACATTGGAGTCCATGATCC
ATTGTTAATTAATTTTTGTATATAGGTAGATGTTTAGGTTTAGGGTTTTT
TTAAAAAAAATTACATATGTTTAAATTGCTCCAGTTCCCTTTTCATTGAAA
AGGGTATCCTTCCCTCATTGAATTGCCTTTGTGAGAAATTAATTGGACAT
ATTTGTGTGAGTCTATTTCTGGGCTCTTTATCATGTTACTTTTAAAAAAT
GCATCAGTTCCTCCACCAATACCTCATTGTCTTGATTATTGCAGTTATAT
AGTAAGCCTTAGCATTAGGAAAAGTGTTTTTCTGCTTTTATTCTTTNTCA
AAAAATTTTGGATATTCTAGGGCTTTACATATAAATTTTAAATAACT
TTGTCTATGTCTAACCGAAAGCCTTATGAAGATTTTGATAAGAATTGCAT
TATGCCTATACATTAATTTAAAAAGAACTGATGTCTTTATTCAAGTTGATT

FIG. 3 (3 of 52)

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CTGCTAATCTATGAACAAGCATCTCT...CAAAGCATTAGTCTTTCTT.
 AATTTCTGTCAATTAATTTTTTAAATTTTCATCCTAAAGATTCTGTATAT
 GTTTTGTGTAATTTATGCTTAAGCATTTCACCTTCTTGGTAACAATTATA
 AATGATTTTGTGTTTTTATTCCACTAGTTCATTTTCAGTGTGTAGAAAA
 GCAATGAATTTTGTGTGTTGATCTTTGTTCCCTACATCTTGCAACATTAT
 TGAACCTCAATTTATTAGTTCCTAGGAGGTTTTTTCATTTTCTTGTAGATAC
 CTTGAGATTTCTATATAGACAGTCATGTTGTCTGCAAACAGGCACAGTT
 TTATTTCTTCTTTTCAATCTATATGCCCTTTTTTTTTTTTTTGGCCTTAT
 TGCAGTGGCTAGAATCTTAGCACTATGTCAAATAGCATTGGTGAAAGCA
 GACATCCTTGTTCCTTGTCTTAGAGGAACATTTGGTCTTTAATCTTGGAT
 TGGC

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GCGCCTCCTTTTCTCTTCCAAAATTTCTCTTGTCTAGTTATTTGTCCAGG
 GAAATTTGAAAGCTCACTTACTGTGCAAGTCAGCAGGAAACAACCTGGGTC
 TGTGCACAGCACCTAGCAAAGTTCTGCTCTAGGAATTACACTTTGGCCCT
 GAGGTAGATTTCTACAAGAACCCTTACCTTCTAAGCAGCACTGGGGTTCAT
 CTTTTCCCAAGTCCTCAGAGCCCATTTTCACTCCTGAGTTCTCCCCACA
 AAGGACATTTCAACGTTGAGTTTATTACTCAACAGAAAATGGAATGAAG
 TCCAAGACCTAAGGAGATAGAAAGGGGACCAGTTATGGCATCTTCTCACC
 CCAGGACACCTTGCTGCATGTCTCTAGTGTGCAACAGACCCTGGCCTTG
 CTCTGTAGTTTGAATGCTCGCTGCAACCAGAAAGGCACCAAGGGGCCAG
 ACCATGCTCTCCTGTCTATCAGCCTTCAAAGCAGAATTTCCCAAACCTT
 GAGTCACAGTGCTAACACACGGGGTGCCATAACATTTTGTGATTGTTGG
 CATTTTACAAAAATAAAATAAAAAAGTTAAAAATGCATTGCTCTATTCTT
 GGGGCTGGCACACTATTGCCCTTTGGCCAAATCCGGTCCCTGACTGTTTTT
 TAAATAAAGTTTTATTGAAACACAACCATGCTCTTGTGTACATATTGTC
 TCTTGGCTGCTTCGAAGCTACAATA

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GTGTTTCGCTTTTTAACACTTACCTAAAATTACTCTGTAATCCATGGATCC
 TTAATTTATTTAAAAAACTAATGTTAATGAGTAGCTTTATTTTCTCCCA
 TCTAATTTAAGGCCACAGAACACCTTCACTTACCTCAATCCTCTCCCAA
 CTTACATGCTTTTAAATGTCATATATGTTAATACCGTATACTTTTAAACT
 TTCTAAAAATAGCATTATTTTATAGCATGAGTGTTCATTTACATTTTGGCA
 TATATTTAGAAATTTTCTTGTCTCTTCGTTTCTTCTTCTATTATGACTCC
 CCTCTGGGATCAATTTTCTTCTACTTGAAGTACATAGTTTAGAACTGCAC
 TATTCAATACAGTAGCCACTAGCCATGTGTAGCTATTGAAGTTTAAACTA
 AGTAAATTTAGTAATATTAAAAACTCAGTTCTTCTCATCTCACTAGCCAC
 ATTTCAAGTGCTCAGCAGCCACGTGCGACTAATGACTACTGTACATCAA
 CATATAGAACATTTCCATCATGGCAAAGAGCTCTATTGATAGTGTTCATC
 CAGAGTTTCTGTTCCAGGACCAAACCTGAGGGTTGGGCTGCTATTTCTCAT
 GGCCCAATAACAAGATGCAGATGAGCTGGGGAGGAAGAGAGTTTTTATTT
 CTGCNACCATTTTACCGGGAGAAGGCCTGGAAATCATCACCAGGCCAACTC
 AAAATTATTACGTTTTTCCAGAGCTTATATACCTTCTAAGCTATATGTCTA
 CGTGTAAAGTGTGCATTACCTGAAGACGTTAGTGATTAACCTCTTTTAAAT
 CTGTAACCTAAGGTCTGAGTCCGGAAGATCTTCCCCTGGAGCCTCAGTAAA
 TTTACTTAATCTAAATGGGTCCAGGTGCTGGGGTAATTACCCTTATCTTG
 TCCCCTGCTAAATCATGGAGGTTTGGGGATTCTTTAGAGCACCAATAAA
 CTTGTTTGTGGAGGCCTGGGGGTTTCTTCTGACCCACAATAAAACTTGT
 TAATCCTAAATGGGTCTGTTAAGAATTCCTTCTTTATTTTGTATATTT
 TAAGGCCCAAGAAAAGCCTGGGCAAAACTCTTGATGGGCTTTTGTACAT
 TCCAGCCTTTGTATAAGAACACTGGTTTTTAAATATTTAACTTAACCATTT
 AGTCAGTACTGAAACAGTTGTTATAGAGATCTGCATTAGTGAGACCTGGC
 CTGCCACATTTCTTTTCTGAAGATCTTATGGTAGTGATCACCTTTGTGA
 AAGGAAAAATAAATCTTGGGACCTCAAAATCACTAAGCCAAAGAAAAAAGT
 CAAGCTGGGAAGAATCTGACACTTAAATCCAACACTGCTAACTCATTCTAT
 CTCCTCATTCTATTCTTTTATTTCTTTTTTCTTTCTTTTTTTTTTTTTT
 TTTTTTGAACGAAGCTTGTCTGTCTGTCACCCAAGCTGGAGTGCAGTGGAT
 CTCAGGTCACTGCAACCTCCACCTCCCGGGTTCAAGCGATTCTCCTACCT
 CAGACTCCTGAGTAGCTGGAATTACAGGCACCTGCCACCACGCCTGGCTA
 ATTTTTATATTTTAGTAGAGACGGGGTTTACCCTGTTTATCAGGCTGG

FIG. 3 (4 of 52)

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TCTCGAACTCCTGACCTCGTGATCCGC...CCCCCTCGGCCTTGTGCT
GAGGTACTGTCTAAATGCTGGAAGTGAAGTGGCAAGCAAGACATCCCTA
CCCTTGAGGAACTGTAATCTAGTCGGAAATACAGATGTCAACCAAGTCT
CACACAAGANATTGTACAAAACCCCTAGGA

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GGAAAAACCTATCACCGCCTCCTATGGAACCTTAAACAAAAAGAAAAGTA
ACAAAGGAAATGAATATTTTCACTTCTGGAAGAACATTGAAAAAGAACAGGA
AGAAAGAGAAAGCACAACTCGAACTGTCCACTAGAATTGACAACACTCTGA
CAGAATGTCTGAACCTCATCGAAGGGTAAAGTGAATAAATAAGCTCCTC
CAGCTTTGGCCCAAAGTCTTATAATTTTTAAACATATTCCTAAATATAAT
ATAGGAGAGATAGCCTTCATCTAAGTAGAAATTTAGCTACTCTTGTAAT
ACAGAGTAATAATAATGACATGCCATAAACAGTGTCTTTGTGTAT
CTGTGCTTTTATAAGCACTTACAAACAAACTGAGGCACAAAGAAGTT
CCACTGTTACTGACCACTTTACAAACAAACTGAGGCACAAAGAAGTT
GGAAAACTAATCCAAACAAACTGGCTCCAAAGGAACCTTGCTTTCTTTG
GGTATCAAGTTCTGAAGAGTACACATTTAACATTGAAACTGAGGTGAGAA
GGCAAGTTTCTATGTAAAGTTGGAGTATTCTGAATACTCTGGGTAGCTAC
AAATAGTATTTAAATTTTATCTTGGAATCTGCAGATAAGGATAAAATAGA
TGGTAGGCAAGAGTATGATCCTTAGGAGAAATTTTTCTGAAGGAAAAA
TATATTAATAAAAAATGATGGAATAAACTTCTAAGATCCTTGCTAGAGC
AAAACTCATTGAGTCTTTGGCTGGTAATGTTGAACATCAACAAAAA
GGAAAAAGTTTCAAGTCTTAAAGTCTACTCCAGGCAACATTTTCAACATCCAG
TTAAATATTAACATTTCTCTTTGTGGAATTGAAGTGTCTTTTCT
TATCCTCTTTTGGTTGTTGTATTATTTAAATGAGTACCTTTTATT
ATTGAAATCATTCAAGTAATGCAGATAAATGATCAGCCCTCTCCCTGTA
CAACATACATACTTAGGCATCCCAACTTCTCTGAGGTGACCACCA
TTGCCAGTCATTCTGTTTTCATGCATGTCCATACAGTATAGGTATG
TCGAGAAATGAAGTATTATTTTTGTGAGTTGCAATCTTTTATTACA
TTTTTGTGTACTTTGGTTGTCTTTCTGTGTTTTCTAGTACCAATGTT
ATGCTGACTTAGGCAGATGAGTTGAGTATTTCTTTTGCCCTATAAAC
TGAAAATAGTTTGTATGACATGAGAATTATTTTATTTTTGAAGGTTT
ATAAAACTTGCCATAAAATCGTCTGGACCGGTTTCTTGAGGATGCCT
GTGTTAGAGCC

>Contig16

CGCTTTAACCTGGGCTACCAATGGTTCGTCAGTTCTAGATTCTCTATTA
ATACCTTTTTCTGTGTCTTTCTCTGGTCTGTTTTCAGCCCCGAGTCTCT
TAGATCTGTCTCTAATATTCCTATTGACTTTACTTCATTTTCTAAGTCT
TTATCCTTTTGCTTTACTTTCCGAGAGACCTGCTTAACCTTATCTCCAA
CTCTTTTATTGAATTTCAATTTCTTTTACTATATTTTTTACTTTGAATA
CACCTCTCTCTTCTCACATTTTCCCCATAGTATTTTGTCTTCAATTGA
CAGTTCTACTATCTTATTACTCTGGAGATATTAATAAGTTTTTAAAT
TATTTTATTTTTATTTTCAAAACAGTGTCTTACTCTGTCACTCAGCTG
GAGTGCAGTGGTGTGATCATGGATCACTGCAGCCTTGATCTCTGAGCTCA
AGCTATCCTCCTGCTTCAGCCTCCCAAGTAGCTGGAACACAGGCATGTG
TCACCATACCCAGCTAATTTTTTGTGTTTGGAGTGGAGTCTCACTCTGT
AGCCCGGTCTGGAGTGCAGTGGTGCAATCTGGGCTCACAGCAACCTCTGC
CTCCTGGGTCTGTTCAAGCAATCTCCTGCCTCAGCCTCCTGAGTAGC
TGGGATTACAGAAACACACTACCATGCCAGCTAATTTTTGTATTTTTGT
AGAGACAGGGTTTACCATGTTGGCCAGGCTGGTCTTGAACCTCTGACCT
TGTGATCTGCCACCTTGGCCTCCAAAGTGCTGGGATTACAGGCGTGAG
CCACTGCACCCGGCCACTAATTTTTAAATTTGTTAATAAGACGAGGTCTT
GCTATGTTGCCAGTATGGTCTTGAACCTCTGGGCTTAAAGTAACTCTCT
GCCTCAGCCTCCCAAGTGTGGGATTACAGGTGTGAGCCACTGAATCTG
ACATTTTTTAAAGTTTTCTCTCTTTACCAAGTCTTTTTCCCCCTTCT
GCTTTTTTGGGTTGTTTTATTTTGTATCTCTATCTTGCTAGAACTTTCTG
CAGACGTTTAGTAATACTAGATTTTTGAGAGTGGGCAACTGGAAAGCTGA
TTGGAACTCTGAATACATGGGTGAGGCTTGTGGCTGTGAGTGTCAATG
CTTGATGTCTGGCAAGGCCAATGGGTTTGGGACCCCTACTATTAGTATA
GGCCTGATCCCTGGGAAAGGCTCTTTGATCTCCTGCCTGGAGGATAAA
GGCCTGGCTACCAAGCCTTCTGTGTGAATGTGAGGGAGAAGGGCTGGAGT

FIG. 3 (5 of 52)

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ATTCACATCATGCTGAA.CCTTTCAA.JATCATCTTGTTTTTAGTAATC
TCCTACCTTAACCTCTCTGTCTTCTGCTAGTATGGGAAAGATGACCTGAAA
ATCTAACCATTTATTTTTCCCCCATTAAATATCATTTTATGATTATTCAGA
AGTTAAATAATTGTCTGCTGTCTCCAAAAAGACTGAATCAACTAGCAA
CAATAAAGAAATTTTCTCAGAGCTCTGCCAGCATTTTAAAAGAAATAGCTTT
ATTGAGCCCAGGAGGTCAAGGCTGCAGTGAGCTGTGATTACACCACTCTA
CCCCAGCCTGGGTGACAGAGCAAAACCTGTCTCAAAAAAGAAATTTAAG
GAACAGCTTTATTGTTGTAAAATAGACATACAATAAACAGAGCACATATT
TAAATTGTGCAACTTATACTTTGATATAACCTGTGAAAACATCACCACA
ATCAAGATAGTGAATATATTTATCACCTCCTGATACAGTTTAGCTCTGTG
TCCCCACCTAAGTCTCATGTTGAATTGTAATCCCCAATGCTGGGGGAGGG
GCTTTGTGGGAGGTGATTGAATTGTGGGGGTGCACTTCCCCCTTGCTGTT
CTTGAGATAGTGAATGAGCTCTCATGAGCTCCCCCTCACTCACTCTCTTT
CCTGCTGCCATGTGAGGATGTGCTTGCCTCTTCTTTGCCCTTCTGCCATG
ATGTGTTTCTGAGTCTCCTAACCATGCCTCCTGTACAGCTTGACAGAA
CTGTGAGTCAGTTAAATCTCTTTTCTTCATAAATTACCCAGTCTCAGGTG
GCTCTTTATAGCAGTGTGAAAAGGAACATAATACCTCCTAAGTTACCTC
AAGCTTGTTTTTAATTCCTTCTCCTCCCTTCTTCATTGCCAAGCAAACA
ACCACCTGTTTTCTGTCACTATAGATTAGTTTACATTTTGTGGGTTTTTT
TTTTTTTGGAGACAAGGTCTGACTCTGTTGCACAGGAGCAGAGCAGCGTA
TC

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CGCGTTATAGGAGATGCGAACTTAAGAAATGATGATAAGGAGACTTTATT
AAATATAATTTTGAATTATTTTGGCATTACAGAAATCTAATTATTTAAA
ATTCTATTTCATAATTTTAAATCACTGTACTTCCAAGCTTAGCTTAGAAT
CCTTCTGTGCTGAGGATTAATTTAATTTGTCTTTATAGGCCTTATCTA
AAATCCAAGAATAATTGCCAGAATCAACCACCTTCTAAATCTGTAAGTAG
AAATTAGTCTTTTTTAAAAATATGCATTATAAGTATGATTAGTAATAAAA
ATAATAAAGATGTTAGCAACCTAAAGAACATGTATTGAAAGGTATTTCT
TACAGATATAAAAAACAGTTTGGTTTAAATAAGAGACAATCATTTTTTGAAA
AGTATGACATTTTTTGAAAAGTAGTTTAGTTTATTAAACCAAGAAAAGCC
TCAAGTGAACTTTAGTCCTCTTGATAGCTAACATTTATTGAATGCTTACT
GTGTGCCTGATACTTTTCTGACTTGCACTACCTCACTGAGTCCTCACAAT
CTTATGAGGCTACTATTAGTAGCCCCACTTTACAGATGAGCAAACCTAAGT
CACAGAAAGGTTAAATAGGTGCTATAGCTATTAAGTGACAAAGCTGAGAG
CCTGTGATCTTAACCACTTTGGTATGCTGCCATGAAGTTAAATAGCTCAG
TAGTCATTAAAAGAGAACATTTGCATTGAACCTTCCAAGCCACTTAACAA
GTATATGCTTCTAATCAATTTAATTTAGCTACATTAGATAGAATGGTAA
AGGATCCTTAACCTTAAAGTTTAAATGGAAGAAATTAGCCCTCTGAAAGAG
GCACAGATTATTCATCTGCAATAAAAAATCTCACCTTTAGTTTTTTAAAC
ATAGTTTTTATCTGTGTTCTGAAATGTAACATAAAACAGTGCTTCTGAA
TGAAAAATTCTCACTGGTGAGAAATTTAATAAGTTTTAATGATTCACCAA
ATCACTTCAGTCATATTTTCACTCATATGCATATGCATATATAGACATATA
AGTTTTTATCTGTGTTCTGAAATGTAACATAAAATAGTGCTTCTGAAAGTG
AAAAATTCTCACTGGTGAGAAATTTAATAAGTTTTAATGATTCACCAAAT
CACTTCAGTCATATTTTCACTCATATGCATATGCATATGTAGACATATATA
TGTTGTATGTATACATGACATCATTAGACACTGTGAAGGATAGCAAAATG
TATATAAGGCAAAATTTATGAACAAATGGTTTAAAGTTTGGGAAGCACTGG
GTTACACTTTTACTTTATGCAGATTGAACCAGTATAGTATGCAAGTCTTA
AGGAAAAATCTACTGGAAGGGCCCTCATTACAGACTTCCAGAGGCTTCT
CTGGAAGTTGACAATACTGACTTCAGTACATCAGCTCGTAAATGAGGATG
ATACCTACCTTATCTGCTTTACACAGTTGTAAAAGTAAAAAGTGAACCTCA
GGAAGGGAATTACAGAATTTAGGAGAACTAAAAGCAGATGTAAATAAT
AGTCATCATTACAGTTATATAATGCTTGACAATTTATATAACACTTTTCA
TACATGACAACAATAACTAACACCCAGACATGTTTATATACATTACCTCA
CTCAGAACAACCATGTGAGGAAGTTGGCCATATGCTTTAATGTCCAAACC
AGGACACTTTTGAGAGTAAAAAGCAGTACTCTTTGACCAACAGGCATAAA
TCAAAACTATCTTGTGAAAACCGGATATATGGCATCCTTCTAGATAAT
AGTACTTTTACTATTATTAATTTTGTCTGTGAATCTAAACCTGCTCTAAA
AAAGTTAATTTTAAAAAGTAATGAAGTACTGATACATGCTACAACATGGG

FIG. 3 (6 of 52)

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TAAATCTTGAAAACGTTAAGCTAAGTG...AGAAGCCAGACAGAAAAGG...
ACATATTACATGATTCCATTTTATATGACACATCTAAAATAGGCACATCTA
TAGACATACAGAGACAGAAAGTAGACTAGCGGTTGCCAAGAAGTGCAGG
AGCAGAAGATGGGGAGTGAAGTCCCAATANGAAAACGCATTACGT
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TGAATCGCAATGATATGTGCCACTTTGCACTCTCTGTGACATATATAATT
ATTTTAAATGCATTCAATTTTTCTCAGAGTGCATTTCGTTTGAAAACATA
GACGGGAAATACTGGTAGTCTTCTGTGTCAGTTAGAAACACCCAAACAAT
GAAAAATGAAAAGTTGCACAAATAGTCTCTAAAAACAATGAAACTATTG
CTGAGGAAATGAAGTTTAAAAAGAAGCACATAAGCAACAACAAGGATAA
TCCTAGAAAACAGTTCTGCTGACTGGGTGATTTCACTTCTCTTTGCTTC
CTCATCTGGATTGGCATATTCTAATATCCCTCCAGAACTATTTTCCCT
GTTTGTACTAACTGTGTATATCATCTGTGTTTGTACATAGACATTAATC
TGCACCTGTGATCATGGTTTTAGAAATCATCAAGCCTAGGTGAGCACCTT
TTAGCTTCTGAGCAATGTGAAATACAACCTTATGAGGATCATCAAATAC
GAATTCATCTGAATGACGCCCTCAATCAAAGTATAATTGAGGCCAATGA
TCAGTACCTCAGGCTGCTGCATTACATAATCTGGATGAAGCAGGTACAT
TAAAATGGCACCAGACATTTCTGTCTCCTCCCTCCTTCACTTACTTA
TTTATTTATTTCAATCTTTCTGCTTGCAAAAAACATACCTCTTCAGAGTT
CTGGGTTGCACAATTTCTCCAGAAATAGCTTGAAACACAGCACCCCCATAA
AAATCCCAAGCCAGGGCAGAAGGTTCAACTAAATCTGGAAGTTCCACAAG
AGAGAAGTTTCTATCTTTGAGAGTAAAGGGTTGTGCACAAAGCTAGCTG
ATGTAACCTCTTTGGTTCTTTGAGACATTCTTACCCTCAATTTTAAAA
CTGAGGAAACTGTGAGACATATTAAATGATTTACTCAGATTTACCCAGAA
GCCAATGAAGAACAACTCACTCTCTTTAAAAAGTCTGTTGATCAAACCTCA
CAAGTAACACCAAAACCAGGAAGATCTTTATTATCTCTGATAACATATTTG
TGAGGCAAAACCTCCAATAAGCTACAAATATGGCTTAAAGGATGAAGTTT
AGTGTCCAAAACTTTTATCACACACATCCAATTTTCATGGCGGACATGT
TTTAGTTTCAACAGTATACATATTTCAAAGGTCCAGAGAGGCAATTTTG
CAATAAACAAGCAAGACTTTTCTGATTGGATGCATTCAGCTAACATGC
TTTCAACTCTACATTTACAAATTATTTGTGTTCTATTTTTCTACTTAAT
ATTATTTCTGCAATTTTCCCAATATTGACATCGTGTATGTATTTGCCATT
TTTAATATCACTAGACAATTCAATCAGGTTGCTACGTTGGTCCCTTGGGT
TTACTCTAAATAGCTTGATTGCAAAATATCTTTGTATATATTATTGTTTTT
TCTCCTATCTTGTAATTTCTTTGAGCACATCCCAAAGAGGAATGCCTAGA
TCAATGGGCACAAATAATTTGACAGCTCTTATTAAACATTATTCTGTAAG
TAAAAACTGAAGTACTTTTCAAGTATCACTAGCAACATATGAGTGTATCAG
CTTCCTAAACCCCTCCATGTTAGGTCAATATGAACCTATGATCTAACAAA
TTACAGGGTCTTATCCCACTAATGAAATTATAAGAGATTCAACACTTATT
CAGCCCCGAAGGATTCATTCAACGTAGAAAATTCTAAGAACATTAAACCA
GTATTTACCTGCCTAGTGAGTGTGGAAGACATTGTGAAGGACACAAAGAT
GTATAGAATTCCATTCCTGACTTCCAGGTATTTACACCATAGGTGGGGAC
CTAACTAC
CATGCACACACAATCTACATCAACACTTGATTTTATACAAATACAATGAA
TTTACTTTCTTTTGGTTCTTCTCTTACCAGTGAAATTTGACATGGGTG
CTTATAAGTCATCAAAGGATGATGCTAAAATTACCGTGATTCTAAGAATC
TCAAAAACCTCAATGTTTGTGACTGCGCAAGAAGAAAACCCCATGCTG
CTGAAAAGTCAGTTGCTCTTTGTCTCCAACCTTACTTCTTTACCTCTCAT
ATGTTTGTGAATAAGCCCAATAAGCAGACNCCTCCTACAAAGTGAACCTG
GTCTCTTCTCCTAACAGG
>Contig19
GTCTTGTAACACAGGTAAGACGAGTTCAAGTTTTATTTCTTGNTTTTAGA
ACGGTAGTGAGCGGTTTTTCAGCNTGAGACCACACCTAAGGTAAGTAGCTG
AATTGGGGTTTTGTCTTGGCTAAAGTTTAAACAACAGCTGGTCTTAATTT
CTCCTTACCATTAGAGCACTCAGTAATCATATAAGTTGTGTGATCATTCA
TTTGGCTTAACTGTTTGTCTGTTTTTATTGCTGTTTCAGTCTTTTTCC
CATTGGGTTTGACCTACTCTATCTGACTTGATCAAATCCAAAGGAAATTT
CCAAATTATGGGAATGAGGCCTCTGAAGTGGCTAAATTTCCACCCCTCCC
ACACACACAAACGTGGTATGGTGGGGGAAAAACGCCAGCAAAAGAAAA
AAAAAAGGAAAGATGTTTCAATTTGACCACCAAACGGGCTTTATTATC

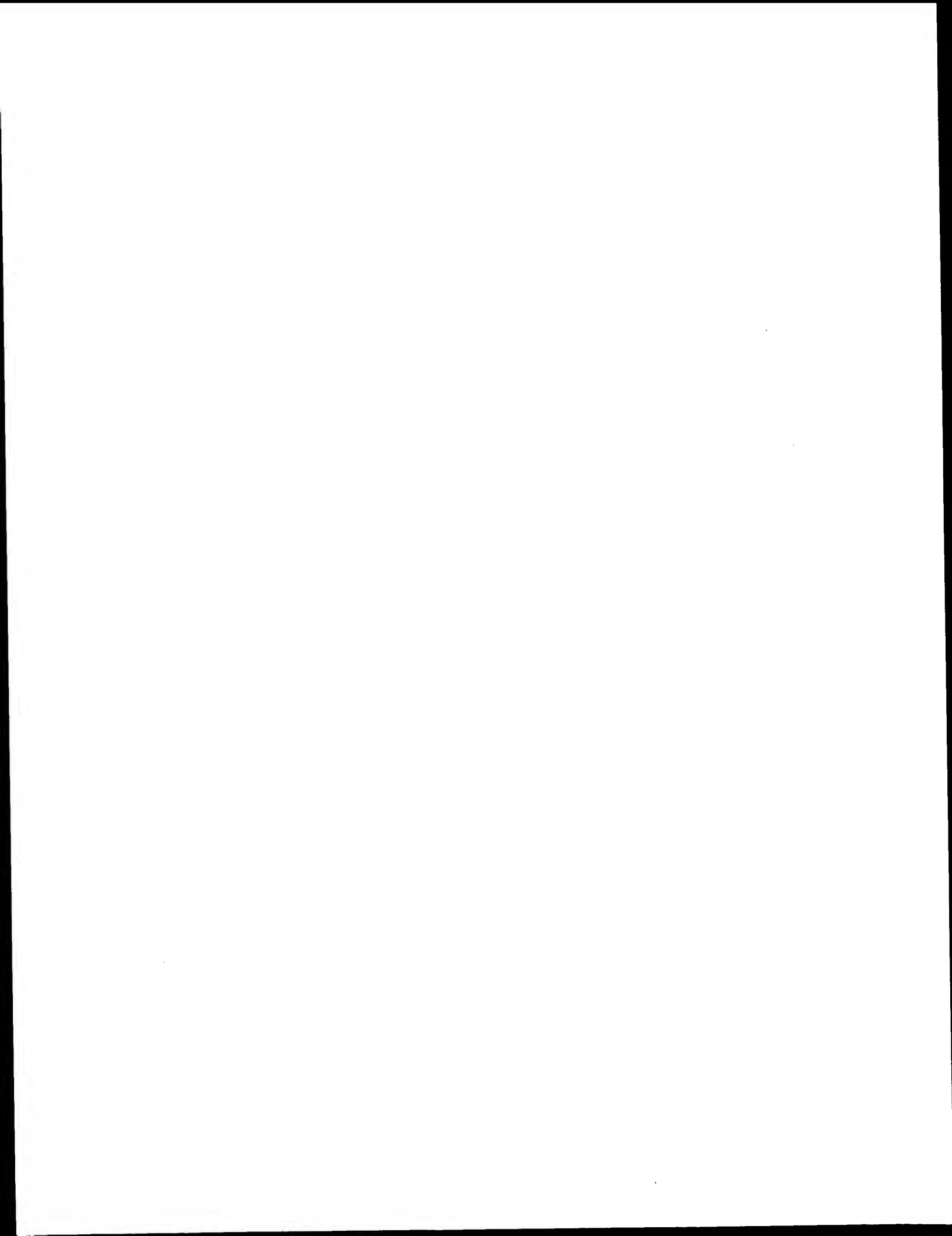
FIG. 3 (7 of 52)

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ATAACAAGGCCACCTTT...GCTAGCCA...CCATACTGAAAGAGCAATGL...
TGTTGCCCCATGCTGTGGGTTCCATAGCTAACGTTCTGCCTTTTTTCTTA
CCACGACAGCCTGGGTTTGGTTCCTAAATCAAGCCTTTTCTGGTTTGATA
CTTGGTAATGCTGAAATAGCAGCAATTTGTCTAGCTGAAATATCGTAAT
AAGATTTTAAAAGATTATTTTAAAGGACCTCAATAGTTAAAAGTCAGCT
TAATTTAAAAGCTAACATCCAAGATGTGTGCATGTGTATGTATGCGTCTTT
GTATTTAAATAGCCCTCATGTTTTTTTTTTTTCTTTCTAGGAACTTGCCTT
TTTTTGAGCAAAAGTTTTTTTCTTCTCTGTTGACTGGATTCTGTTTTCTT
CATTTACTTCTGCTGTCTCTCTTTCTCTTGCACCGTCTGCTGCATGAGA
GCCCTAAAATAGTTTATAATAGCCTGGGGTTCTTAAAGAAAATGGAGAA
GGTGCCAGGCTCCCTTTTAGGGAGAACTTCTATTTTTCTTATGGAATC
CCTAGAGTGTAAACAGACAAGTTTCAATTCAGCTCTTAACTGCTTGCCTT
TGTGTTGTGTTACCTGATTTTTTTGACTATTATATTTTTGACTAGCTATT
GCAACAGAAGCTACTCTTGGGTTTTCAAGGAAGATTGTAGTTTAGACATG
TAGAAATGTCTTTAAAAAACAACCTTTTTTTAAGTGCCTGTAA
AAGCATCATATGGTCTAGCCTCCTAATAATTTTCCCTTTTTGGAGACCAG
GATTCAGGGTGGGCTCTGCCCAGAGCTCAGAGATCCAGTTAAAAGAGAGG
TAGTCTCGGCCGGGCGTAGAGGCCAGCCTGTAATCCCAGCACTTTGGGA
GGCCGAGGCGGGCGGATCACGAGGTCAGGAGATCGAGACCATCCTGGCCA
ACATGGTGAAACCCCGTCTCTACTAAAAATACAAAATTAGCTGGGTGTG
GTGGCAGGTGCCCTGATGCTCCAGCCACTCGGGAGACTGAGGAAAGAGGAG
AATCGTTTGAACCCGGGAGGCGGAGCTTGCAGTGAGACGAGATGGCGCCA
CTGCACTCCAGCCTGGCGACAGTGAGACTCCGTCTCAAAAAAAAAAAGAT
AGGTAGACTCGATGTTGTCTGACCCGAGCAAGTTAGAGCAACGCCACACT
TTGAGACGAATTTAAGAGTCTTTATCAGCCGGCGACCAAGAGACGGCTA
ACGCTCGAAATCTCTCGGCCCTTGGAAAGGGGCTTGATTTTCTTTATG
CTTTGGTTTAGGAAGGGGAGGGGAGCTCAGTTGCAACAATCTACAGGAG
TAAAAACATGCAAGAAATTA AAAAGACAAGTGGTTACAGGAAAACAAC
AGTTCAGGTGCAGGGGCTCTAAATCTATCATAAGATGTTAGGTATGGGG
GCTCTGCCGGACACAACTCAAGGCTTTATGCTGTTATCTCTTGAGCGAA
ATCCTGGGAACCTCGTACATTGCTTGCTTCAGTACCTTATCAGTTAATCG
GACTCTTTGATATGTTGGGAGTCAGCGTACACAAGTTAACTCCTTGAGGA
AGGGGGTGGGTAAGGAGTCCTTGATGTCTGGTAAATGAAGGAGCGAAATC
GAGTTCTCTGGCTTTCTCAGCTAAGGGAGAGCTTATTCATGTGGAAACA
AGGCTAAGTGATTAAGGGAGAAAGGGAGAGTCTGAAAACAAGGTTAGGTA
TTACAATGTCAATAAAATTTGGTCTCCTTATACAGTCTTATGGTAGATTTT
TTTCCATCTTTAATCTCCCTCTAGCACCACCAGACTTTTTCTCTCTGTAC
CTTGAGATGTAAATTTTGCTATCTGAATTTTCTGCTAAGAGTTGTTTCT
TTAATATGCAAATTTAGGGTTATTTAGCTGACAACTGCCAAAGTAGTGAA
ACAAGTTATCAAGAACTTGAACGTCTAAGGTAGGAAAAAAAAAAGTCTTT
ATGAATCTATAAGATGTACTTCTATTGGCATGCCTAATACGTCTATGTAT
TTACGTGTTGTGTACACAGTTTTTCACTACTGAAAATATATAGAGGAGTT
CTAATTAATTGACTTAAGACAATAAAAGCGCTTGAATCAAATACCTTATC
AGGAAAAAGGAAAAGACAAGTCAAATGCTTGTTCAGTCTATATACTTA
AGTAAATCTTTAATAAATAAGCTAGCTTTAACATTATTTGAAATGTCTT
AAGAATTGCCAGCAGGTTCTGGGTTACAGAATAGTGGGGGTGCAGTGGG
GTGAGGGTGGTGGGGTGGGNGGTNNNACNNNNNCNCCCCCCCCCCCCC
CCCCCCCCCCCCCTCCCCCCCCGCGCCGCGGGCGCGCCCCCCCCCGC
CCCCCGCCCCCCCCCGCGCCCCCCCCACCCCCCCCCCCCCCCCCCGC
GCCCCGCCCCCCCCCGCGCCCCCCCCACCCCCCCCCCCCCCCCCCGC
CCCCCCCCCCCCCCCCCCCCACCGGCCACACGACCCCCCCCCCCCCGAC
GCCCCGCCCCCCCCCCCCCGCAGCCGACGCCCCCCCCCCCCCGCCCCG
CCCCGACCCCCGACCCCCCCCCCGCGCCCCGCCCCGCCCCCCCCCCCCG
GCCCCCCCCCCCCCGCGCGCGCGCCCCACCCCCCCCCCCCCAGCCCCGACC
GCGCGCCCCCCCCACCCCCCCCCCAGCCCCCGCCCCCGCCCCGACCC
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GGCAGTACGCTATAATTCCTCTTCACCTTACCTCATCTGTTCTCTGATG
GATGACTTTTTTTTTTAGTTTTCTAAATTCCTTTTCTTTGCTCTGGAG
ATGGGTGATTGATGTAGTCTGGGTATTTGTTCCCTCAAATCTCATGTTG
AAATGTAAATCCCCAGTGTTGGAGGTAGGGCCTGGTGGGAGGTGTTTGGAT

FIG. 3 (8 of 52)

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CATGGGGGCAGATCCC. ATGAATAG. GGTACTGTCTCTCAATAG. 3
AATGAGTTCTCCTGAGATATGGTTGTTTAAAAGTGTGTGGCACTCCCCCA
TTGCTCTCTTGTACTGCTTTTCGACATGTGACATCCCTGCTCCCTTCGC
TCTCTGCCATGATTGAAAGTTTCTTAAGGCTTCGCCAAAAGCTGAGCAGA
TGTGGGTGCCATGCTTGTACAGCCTGCAGAACTGTGAGCCAAAATAAACT
TCATTTCCATATAAAATTACCCAGCCTCAGATATTTCTTTATAGCAACATA
AGAGTGGCTTAATACAGGCTGGGCATGGTGGCTCACGCCTGTAATCCCAG
CCTGTGGGAGGCTGAGGGGGGTGGAACATGAGGTGAGGAGATTGAGACC
ACCGGCTAACACGGTGAAACTCCATCTCTACTAAAAATACAAAAAATTAG
TCGGGCGTGGTGGTGGGCGCCTGTAGTCCCAGCTACTCTGGAGGCTGAGG
CAGGAGAATGGCATGAACCCGGAAGCGGAGCTTGCACTGAGCCGAGATT
GCACCACTGCCTCCAGCCTGGGCGACAAGAGTGAAACTCCATTTAAAAA
GAAAAACAAAATTTCAAACAGAACAAAATGAAAAAATAACCAAGTGAAA
GGCCCCCTATAAAAAACCCCTCTGGGGCCCATCTCCACCCCTCAAGTGA
AACCACATTTAACAATTTGGTGCATATCTTTCAAACCTTTTGTGTACA
CATATAAAAAACATACATGCTTTGATTTGGCTCAGACTGTACATAGTGTT
TTCCCTCTTGCACTTTTACACTTAATATATCTTTGACATCTTTCTATGTCA
GTGCATGTTGGCTCGATGATATTCTATCATTAAATACCCCTCCAAAAATG
GTAAAAATCATTTTAAAAATCATTACACAAAGTACATATTTACAATTTTA
AAAGAAAAACAGAAATCCCAAAACACAACGACAAACCTCTAAAAATAATCTC
TATCTTTCCACCAGCATGGAACAGTTCACTCTTTTTCACATAAAACGAA
TTATGTGATTGGAAAGATTAACCTCTAATCTACACATTTATATACAGAATG
TCTATTTGTTAAGCCTATCTGAAAATAAAAAATTCAGATGATTAATTCA
CTTACACTTAGAAATTAAGTCAATATACTATGAATACACATTGTGATCAG
TTATAATATGATGCTTCTTAGTCTAGGGTTTCAATTAAATAACAGTAAAA
AAAATTGGATAAAATAAGACAGCTAATAACTGAAAAATCCAGAAATTCAAA
GATTATATTGCCAACTAAAACACTGCCATTTACATTTTTTTTCTACTT
GGTAGCAAATGCTAATGGAATTCAATCTGATTACTTAAAGTCAGTTCAC
ATCACACATTCAATCAGGATAATACGAACATAATATGCCTACTATAGCGT
TAGATTAAGACATAAAATTTTTTGGCTTGAAAGTAATGACTGCGTACCAC
TTGAGACATTTGTCAACCACTTCAGCACATTGTTTACGAGTGAAGTGGATG
TCCACAAGGAATAAAACGACAGCAATATTTCTATCCATACAGATTTTGC
AAAGCTTCTCCTCTTGCAAGGTGTCTTAGCTGCTCTTCAGTACTAATCTCT
TTCTGCAATGAAGTCTGACTTGATTCTGCTTGTGTACTGTCTTTCTGAGC
CTTCACTGGATCTGCAATCAGAACCTCAAGTGATTTACAGTTGCTCCCAG
ATGTCTGAATTTTTTCTCCATTATTTTCTTAATGTCTTTGAAACTGAAC
CCCATTTCATATAGCTTCTTGTACCATAGGATTATGGAAGATGGTATCAAT
TTTTCTAGTTAGTGATGGCGTTTTTTTTCAGCAGTTCTTACCAGACACTCT
TAAGTGAATGGGATAAATGAATATTGTTTATATATTTTCGTGTCTTCTGT
CTTAACAGATATTTACACCCTGGATGCCATTAAACATGTTGTCCCAAGGT
CTTNCCTGGGCT

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CTTTCTCCCTTTTTTACCCCAATTTTCGTAGGGATTGGTTAAAACCCATG
TAAAAATCCAAACACCGGCGGGGAACGGGGTTCAAGCTCGTATCCCCA
CCACTTTGGGAACCCAAGGTGGCAGGATTGTGGAAGCCAGGCATTTGAG
CCCACCTTGGGAAAAAAGAGAACCCCAATTTTTTTTGAACAAAAACC
CCAACCTCCAGGAAGAAATAAGTATGGCTGGGTGAAAGTCACCAAAG
ATGGCCGACTGGCTGGTCAAGTAACTTTACCTGATGGTTCGTAGAATATT
TACCTTCACCCAGGTGGGAGAATTGCTTGAGCCAACCTCAGTGTGGATT
CAGGAACCTGATTTAATTGGTATCGTGATTGTGGATTAGATTCTCAGGGA
TGCATTCACTAAGTAAAAGTGATAATAGCTACTTTTAAAGTAAAATAATGA
ATGAATCAAACACTCTAAATCCATGGTGCTATGCTAAGCTCTTTCTGTAT
TTTATCTCATTTGATATTACAAATATTTGATGTGTTAATAGTAATGACTA
TCTCCATTTTTTACAAGTAAGGAACTGACATTGAGAGATTAAAAGACTAG
CACAAATCACAAGTAAATGAGATTTGAATCCGGTCTTGATTCCAAACTC
TACAGTATTCTAAATTCAAGGAGACTAAATTATAAGATGGAGAGCCAATT
TTACTTTATAACAGGTTAGAATGGCAGAAGAGACCTGACATTCACACCT
CTAGCCAGTGCATCATCTTCTGTAGGCAAATATGCAGGAAATCTATAAT
AAGAACGTCCTTTGGTGAAGGCCAGGTGCAGGGGCTTACACTTGTAATT
CAGCACTTTGGGAGGTCAAGGTGGGAGGGTCTCTTGATGACAGGAGTTTG

FIG. 3 (9 of 52)

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AGAACAGCCTGGGCAACAAGTGAGACCTGTCTCTACAAACAAAACAA
ACACAAAACAACTTCAAGAAAACCTCTTTGGTATGGATCAGAACAGATG
AATTATCTATCTGATCCAAATGCTTAATGACATTAAGCCACAGTCCACTC
ACTGCCACAATAGAGATATACCTGCCAATGCCACTCAGGTAATCCCATCA
AAAGTGGTAATGAGGTCTGCAGCATGACTTGTCTTAGTGATCCAGCCT
GAGACCTTGAGATTGCAGCATTTTATTCTACATATGCACAAAACATCTGT
TGAAAAATCTTCTAAATTGATGCAATACATTCTGTATCAAGAATACCTGTC
TGTAATCTCCATAAACCTCTCCTTTCTGTTTTAAAAAATAGTAACAGCA
TTTCTCCTTACATGACAAAGAAATGACTTCACCATCTACGAAATAGTGAA
TAGGAGCTGTGTGGAAGGAAATTAGCTCTACTTCTTGGTGGAGATGAGAA
GGGAGTGTCTCTGAAAATCAAGGCTCTTGTCTAGTGGAGCCAAAGT
CGTTTTTTAGAGTGTGGACAGTTGAGAAGATAAGACAGGGACCATCCACT
CATGTTTTCTTATTCCATAGGCCTCTCTCAATTGGGCAAAGCACTCCAG
ACCTTTTGGGAAGAGTGACACCAAAGGCAAGCACCTGCTTGGCAGGCCCCCT
CAGCTTCTACGCAAGTATAAGTGAGTATATAAAATGGGGTACTTGTGCT
GTTGAGTACCTTATTTCCAAATGAGGCTGCGGGTGTCCCTGTGGCTGTG
AGAAGGCCCTCTACTGGATAGGTGGAAGTTGTGTCTCATCTTTCTAA
CCCTGGATTGACTTGCCCAAAGGAAGCCATTATTAACACTATAATAAAA
CCATCCTTAATCTGGGACTCTCTTCATGCAGTGGTTCTTAACCAAGTGATA
AACATGAGAGTTACTTTTGGAGCTTAAAAAAATTAAGATGCTCAAGGTCT
ACCCAACTGACTGAATCTCCAGAGGTGAGGCCAGGGATGTATACTTTT
GAGCCAGACCTCAGTTTACCCTGCAGAGCTCATAAGGTTGCATAACACCC
TTTGTGAGCCACTCTGATGAAAAGAAAAATTTGGTGAGGAATAAGTTTTAG
AGAAGAAGGAGCAAAGGTGTTCTTGGCCAGTGAGAGCCAATGACAGGGAA
ATGCAAACAATGTATCCACAAGAAAGGTAAATTACCCTATAGAGCATTTT
AGGATAAATGAACATCTCATGCTAGGGTTGAGAGAGGGTACAAAAAAA
AAAAAAAAGACCCTCTGGATACACAACGCGATAAATGGAATAAAGAA
TTTTTCTCTGTAAATTAATAAAATCCTTTGTTACTGAGGTATAATTTAA
TCTATTTTATGTATAGTTCAATGAGGTGTTATAGATAATAAATTTTTTT
GTAAATTATTATATGTCTATATACTCATACATTCATTTTTAAAGTCAGA
AATGTATATAACCATTAACCTTATAAATCATTCAAGTCATTAGAGATATA
GATACAGGAGCATATTTTATATCCACCACAATAATTATTACCATCTCAAC
AATTCATCACCCTCAAATTTCAAGCGTAGGGGTTTTTAAATGTCAAAG
GAGTCTACTCAGTGGGAAGAAAGTTAAGGAAAAACCTTTGGGGCTTTGG
GCTCCTTCCCCCTGGGGTTAAAAAGGCAGGAAATTGGGCTTACCCCCCT
GAAATTGGGAACGAAATTTTGGGAAGTTTAAAAA

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TCAAGCAGCCTTCTTCTTGGCTTCCCAAATTGTTGGGATTACAGGCAT
GAGTCAGGATTCTGGCTTAGTTTACATTTCTAGAGTTTGTATAAATG
GAAACATACAGAATGTATTTTTCGGAGTGGGGAGTGTCTTATTTCT
TTTCTTTTCCATTTTCCCCCCCCNCCCCCGAGACGGAGTCTCGCTCTG
TCTGTTGCCAGGCTGGAGTGCAGTGGTGCGATCTCGGCTCACCGCAAGC
TCCACCTCCCGGGTTCAAGCAATCTCCTGCCTCAGCCTCCTGAGTAGCT
GGGATTACAGGCGCCGCCACCACACCTGGCTAATTTTTTTGTATTTT
GGTAGAGACGGGGTTTACCATGTTAGCCAGGATGGTCTCGATCTCCTGA
CCTCGTGATCTGCCCGCTTCGGCCTCCCTAAGTGCTGGGATTACAGGCGT
GAGCCACCGTGCCCCGCCCAAGTGTCTTATTTCTTAACCAAGCTTTCTG
CAATCTTTTTTATTTTACCATCTCTGTGATCCCACTCCCAAAGGTACTA
GATGTCGATTGGTCTTAGGATCAGCTACCATTTGCCCAACTGCTTTCCA
GCCTTCCAAAAATTTTTTCTTTTTTCTTAAAGATACTCCTGTGTGAGG
CTCAGAACTCTGAATTGCTACTGCAAATATGAACCTCGGTGATGTGAATG
CCAGGGAATTGCCTGATTGATCAAAGAAATGTATCCCCTTCTCCCTCACT
CTTGCTGTCTCTCATTTGTTTTCCCATCCTTGTGGATTCTGTAATTTA
AATATCCCTTAAATGTTATAATATTTAATGGCGTTTGGCGAAAAGTACA
GAATTAGGTGCAAGAGTGATAGCTGTTATTTTTTTTTTGGCCTCTGAGA
CTGTTTATATATGCAAGTTATTTAACAGAAAGTTCTGCAGTGACCTGAGA
TGTCAGGGGGGTCTGATAGAGTACGTTTGAAGGCAGTTACTGGAAAAAA
TAATGCCATTTCTGGTTTGTACTTCCGTAAGTTCAAGTACCCCAATATAT
TGTTTTACATGTGGCATTGAGTAAAAAGTAGCTTCCCCCTCCTTTCTCT
TCCTTTTCTCCTTTCTGCTTCTATAAAGCATCTGCTTTGGGAACTTCT

FIG. 3 (10 of 52)

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TAGGAGGAGAGCTTGCCAGCCCGTGGC AATGGAGAGGTCTTGCAAGA.
 AAAAGAGATGCTCCCACTCAATGCAGGATGGTGTGGAGGTAAATGGGGAT
 ACGTCTGGCATCACTCAGGAATGGGCCTTCTTGGCAGGGAAAAAAGGGA
 GGGGAAAGAGGAAGGGAATTNNANATNAATTGCTGAATACGGGGATTCC
 ATGGCCTGGATCCAGGAAGAGAACTTTGGGAGGTGTGAACCTGGAAGGCA
 TCANCTGATGAGGAGCAGCCTGAACTCCGGGGAGGACCTGTTTTTGGTGG
 CCGGAAAAAAATGCCTTCCACACACAGGGAGGCCACCCGGCTGATGGGC
 TGGGGGTGGACGGACAGCCCTAGGACAGGCTTGGGAAACCAGGCTCAGG
 TAGGGCCTGCGAGGTTCTCGCTGCGTCTCTTCTCTCTGGTCTTAGAAA
 ATAGAATCCAAGGCCTCTTGAGAGTGAAGGTGGGTGGGAGGAGGGCAG
 ATGGGGCTTAGGCCCAGGACACCCGTAGAGCTACTGCCAGCTGTCTCTC
 AGGGACTCTGCTGAGGTCACTCCAAGGATCATTCTTAGCCTTGCTAGACA
 GTACTGACAGAGGGAACCGTAGTATCGCACCCACTTCTTCTCTTCAAT
 GAAAGTTTAAAGGTCACCATTTCTCTGGCAAAGGAAGTTCCACAAATAT
 TCCATTTCCGGTCTTAGAAAACAGCAAGGTATCAAGCAATTGCAAACCTCC
 TGTGCTGGGGAATTCCAAGGAAGTAGGGGCAGAGTTCTGGTGGAGACAA
 AGTGAATTCGAGTGATTAGTCAGTAGCAGTAGCAGTAGCAGTAGCAGTA
 GCAGTAGCAGTAGCAGTAGCAGTAGCAGTAGCAGTAGCAGCAGCAGAACC
 AGAATTTCCCGCACGTGTCTCAGGCTCTCATTTGCCAAGTCAAGTCTCTA
 AGTATTTTTATTGGCAGGAAAAATAAATAGCTATGAGTGAAATAATTCA
 TTAGACCTGAGCCTCCATCAATTTTGTGTTTAAAGGCCTGACTCTCTTTA
 CCTTCTCCTGGGATGGAAGATGCAATGTTCTGATCTCACTGTCAAAAA
 AGAAGAACCAGTGGGTATATTGTATGCTTGAGTTCCAGCCATTAGTCACA
 AGACATAGAGATGACTGCCATGTGTGTAGACTTTCTATAGACTGTGTGCT
 AAACCCGACCTGCCACTTCCAAGGAGTAGATGAGGAATGTCCATGGTTCT
 GGGGAGCCCTACCCCAATTTGGGGCAGACATTCCAAGCTCATTCTCTGT
 GGAGGGGGTTGATGGTTAAAGGAACGGCTGGGATTTACTCTCTCTCTAG
 GGCCAAGAAAAATGACATGCTGCCTCCATGTTAATCATCCTTCCCCCTGT
 TAATAACTATGGCTTTAAGTCCCCGGTTAGGGCCTTCTTCCAAAATTGGG
 GAAAAAATTCCCTCCCCCTTAAAAATTTTTTTTTTAAAAAAACCTTT
 TTTTTTGGGGGTGGGAAAAAAACCAAAAATTTTTTTTCCCCAGGGGTTT
 TTTAATTTAAATTTCTCCCCAAAAATTTGTTTTTTTTTTTCCGCGAAAAA
 AAGACCCCCCAAAAAAAGTTTTTTGGCGGAAAAAATATTTTTT
 TTTGTGTTAAGAAATGGAGAAGAAGGGGGTTTTTTTTTCTTCTCCCCC
 CACCCGCCAAAGGAAAGTTGTTTACAGATTGTTTTGTGTCTCCCCCCA
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ATGTGCCTGCGAAATCATCCTTCCAGAAATATTTGCCCTTTCTTTTGT
 ATAGAGTGGCACTGCCCTATATGGTGACCACTTGCCACATGTGGCTGTG
 AACACTTGAAATTGGCTTGTGAGAAATGCAGTGTAAGTGTAACACAT
 ACCAAATTTCAAAGACATGGCACATAATAAAAAATGTAAATATCTCATT
 AACAATTTTTATATTGACTGTGTGAAGTAACATTTTGAATATATTGGATTA
 AATACATGGATGATGCCCCAACACCCACAGTCCCTTATCAAGTCTCTACT
 TCACATTTTTGTACTTCTGACTTAGAAATAGCACTGGCGTCTAAGAGCCT
 ATTAATGTCGTCAATAGGTTCTTGGGAACCACAATTTTAAACAAAATGAC
 ATATAAGAAAACGAATAACATTGAACAAAATGACATTATTGAGGACCTG
 CTGCATGTTGTTTTCACTTAAAGTCAGTGTCCAAGAACTATCAGTGACAT
 TTAGTGAGGAATTGCTGTCCTTCTGTTTACAGGAACCTGGGCAAGTTAC
 TTAATTCCTCTAAGCCCGGTTTATATCCCTGCAAAGAGAGAAGGATAATA
 ATCACCAGTACTTAGTGATGTGTAAGGAGAAAAATAAATAAATATG
 AAATGGCTGACAGTGTCTTGTGACACAGAAGATGTGTGATCCACAGTAG
 CTGCTATTGTCTGCCTCACTTCACTAGTAATGGTCCAGGGAGGCCTTTAA
 TGTGCATGGTGCAGTACATTACATGTTGGACATGGGTGAAGGGAAAGAC
 CAGGCTCATCTAAACACAATAGGATGCTTGTGGTGTGTTTGGAGGGAATC
 AAGGACTAGTTATCCACAGCTGTAACATGCATGGATCAAAAGAGATAAGG
 CACACAAAAGACTTTGTGATAGCAAAGCATTACAAAATGCAGAGACCAG
 CTGTGGGTGGTGGTGAAGTCAAGCCAGCTTCCCTCTGTGCCTGGCTGAGT
 GGTTCTGGGCAAGTCAAGCCATCTGTCTTGATGCCCTTCCCATCTATAG
 AGAGGGAGCAACTGAGGCCCTTCCAATACTGAAGTCCTTTATTTCTGCT
 ACTTTAGAAATATCCACATTTTGGTAAATTCAAATGATCCAATGATTCC

FIG. 3 (11 of 52)

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ATTTCTTAATGTTCAAAAAGAGCCCCAACATCTAAATGAATCAAAACA
AATAAAATATTTATTGTGTATGTTTTGATTGCTGAACTTCTATTTTAGC
AACACACACACACACACAGAACCCATAAGCCTTCATCTTCTTGGAT
AAACGAGCCTTCTGTCTGGCCATTAAAGTCAGGATTAAGTAAATGATT
CCAACTCGCCTTTTGCAGCAGTTTCTGATGGGTCTTCTGCGTGGCAGTG
GCCCTCCTGACTTATGATTCTCTGTGTGTCGGCCTGTTACCACTGCAGCT
TAACTGAGGAAACAAGAACAAGCCTCTGACCCCAAGAGACTGTTGG
AGGCAAAGGCTTCAGTCCCAAGAACCTCACACGTGGGGAGCCCGAGAGCC
CAGCCCTGACCTTTTCTCCAGTAATAACATAAGAAACAACAGGCCTGGC
CTTATTTTGGATACAAAGAGTGGTGCTTTTCTTAAATCTTCTTTAGTC
AGGGGTACCCCTTCATGGACGCCCAACATCCATGGTTCTGCTTGAGTC
CCTGCTTCCATATTCTGCACTTCTCACTTGAATATCCCTGGAGTACGT
TAAGCAGCCAGGTTTGGAGGTTCTTGCTGTGCAGGCGGTGTGTGCATGT
CCTCTCTCTCAACAGGACACAAGCTCCCCAAATCAGACGGTATGCCTCCA
CGCCCTTCCCAAGCCTCCCAAGCAGCAGCAGCAGTGTGAGGGGAGCTGG
GGCCAGGCCATGATGGGAAGCACTCTCTGCCTAAAGACTAGGGTGATGC
GCCCTCAACTGTGGGAATGAGCCCCAGCTCTGGTGTCTGCCTCGGTTTTT
CCTCCTGGACAATCAACATGAACTCCTCACCCCTCTTATCCACTTTGCAT
AAACTGAAAATAACAAACCCAGGGTCTTTCTGTCAAGGAAAGGGTTTTT
TTTTATAAGATTAAACAGAGATGATTCAACACACCCAGGATATAACACAT
GGGCCATGAGTCAAGGCCAGGCATTGCTCTGGTCAGCCTGTTGTTGGGC
CCCCCTGGCAGGGCTCTCCCTGAATCTTCCCCCTCTTGAATCCCCATCA
CCACAGCACGTCCAGCTTTGGGTACAAGGCCAGTAAATGGGAAGGGGT
CAGATGACATAAAGAGCCCTTTCTGTCCCATGAAATATATTTGGATAA
CAGATGGCATTTCCTGCTGTCTTGGCCAGGGCCAGAGCCTCCACTTG
CTAGAGGCAGACAGAGGATGGAGAGCCCTTCATTAGTGGGAGGACATCA
CAGGTGGGCAAGAAACCACAAGCTTGCCTGAGGCCAGCCTTGAATAG
CAGCACCTGCCGGCACCTGTGGTCTGGGGACAGGGTCACAGGATGGAGGG
GCCTCCTAAGCCTTTTATCTCTATGTACTAAGTACAACCCATTTCCAC
CTCACAGAGCCAGATCAGCCTCTGTGAGGTCTGGTGGCAAAGGATAAT
TGCTTGCCCGCCTGCCCGCGGTGGGGTGTCTGTGCTTGCATTCTGGGA
GGTTGTGGGTACTCTGCAATAGGTCTCTCTGACCAGCTCACCCCTCTA
CTGCAAACTCAAACCACTTCAAAGAAGATCCAGCACC

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CGCGTAGTCTAAAGACTGAGTCTGAAGCTGTCCCTTCTGCTATGGACTT
CAGATTTTAGCCCACTTGAATTGCTCCATATCCTCCAAGCCATGGCCATC
CCTTGACTCTCTGGGCTCCCAAGCACTTGCTGCCTTCATCACAGTTTG
AGTTAAGGCAGAAAGACTGGTTTCCATGTACACTTGTGGAAGCTTCTC
ATTTCTTTATATAATCTCTGTCTTGTCTACTGCTTTAAATCTAGAAA
TTGTTTACAAACACAAAGGTGATCCTTTAAAGCTCAAAGCTGATTGTGT
CACCAATATATACCACTCTTAATGGCTTCCCATTAACCTTTGAGTAAAGA
CTTTATGGAGCCTACATAAGGCCATGACTACCTGGCTCTTATTTCTCTC
TCATCCTCATCTCACTCACTCTCCACTCCTATACCCCTCACTCCTT
CCCCCTCCTCTCTGAGCTCCAGACTCCCAATTACCTACTTCCACCTT
TTTGACCCCAAGGACTTATCTCAGCCTGGAATTTCCCTCTTGTCTC
CACTGAACCTGTCCACTCCAGTCTAAGACATGTGCTTATGTACACGCCC
TTACCGTGCTTATCTCAGTTTGTAATTATCTACTCATTAGAAAAGTGT
GATGAAGGTCTTCACTGTGAGCTTTCAGGATAGCAGGAATCATAGCTGAT
TTTACTTACTTAACGGGGTTTCATTCTTTGTAACCTTTTTTTTTTTGAG
ATGGAGACTCACTCTTGGCCAGGCTGGAGTGCAATGGCATGATCTCGGT
CACTGCAACCTCCACCTCCTGGGTTCAAGTGATTCTCCTGCTTCAGCCTC
CCGAGTAGCTGGGATTACAGATGCCTGTACCACGCCAGCTAATTTTTT
GTATTTTTTGTAAAGACGGGGTTTCATCATGTTGGCCAGGCTGGTCTCGA
TCTCCTGACCTCAGGCGATCCACCCACCTCAGCCTCCCAAGTGCTGTGA
TTACAGGCATGAGCCACGGCACCCAGCCACTCTTTTTTACTTATGGGTG
AGAAGCCATTAGAGATCATTTCTTTCTTTCTCTCTTCACTAAGGCA
CCAGGGTCACTAAGTAGTAGGATACTTTGAACTAGAAGTCAAGAAATTGA
GTTTTAATTTTACCTCACTCTCATATGAATTCTCATGTGACCTCGGG
CCATACTTCCCTGTACCCTGTTTCTCTTTTATAAAAGTAAGAGTTTAA
ACTAGATGGTCTCCGACATGCATCCTTCTCTAACATATTCTGGAACCTTC

FIG. 3 (12 of 52)

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AATAAACTAAGATAAAC AGAATAATTAAAACTTAATTTAAAAGAACA
 GGAAAGGAAGCAGTTACATTAAAGCAAAAGAGACATCTTCATGGTTGAAGA
 AGTGTATGCCCTGGTGTCTGGATCCCATTTAGGAACTTGGTAACCTTGC
 AATCTTGGGCAGATTGCTTAATTTCTCTAGACCATGACTTCTCTTCTGT
 AAGATGTGATAAGAACATCTACCTCACAGGTTTCATGAGAGGATTAATG
 AGATAATGTATTATAATCCCTTGAACATGGTAGGCTGTTATGTTAAGTCC
 TTTCTCTCTCTCTGTAGCTATCATGGAATTTAAAAACACATTATAACTA
 GAGCATGAGTTGCGACTAAAGGCTCAATTGTCTCTGCATGTGTTGGCTCA
 TGCATGCTTTATTCCTCTGAAGAGCTTTTATACCAAGTGAAAGGAAATAA
 TTGCATTTCCCTGAAAATTACAGGAAAAAGTTATGTTTTTCTCTTCATT
 CAAGTGATTCTGTTAGACCCAACCACATGCAACAATTTTAAAGTTGCTTC
 CAAATATATTACAAATATTTCTGTCTTCAAGGAACAATGGCAAGACCA
 TGAATCAGGTTTACATCCGGATTCCACCACTAACCATGTACCCAATTACT
 TCAGTCACCTTCATTCAAGTCTTACATATCACAGAATAAAATCAGATTTT
 ATCAGAGGAGGTGAAGACAGGGAGAGGAGATATTTCAATCCCTTCTCCGC
 AACCCCGTTTTTTTTTTTTTTTTTAAACAAGGATCCTAGAGTTACTGAATG
 ATAGCACGTTTGAGGGGGAAAGACCCCTAAGGATGATCTTTATAAGCCATC
 ACTTGGTGTGGTGGTGATAAAAACTCGAGTATCTTTATGCAGTGGAAA
 GAGAAGATTGGAATCGGAATCAGAAGCTTGAGTTCAAGCACTGGTTTCAT
 CAGTCTTGTGATCTTGGGTTGGTCACTTAACCTCTTCAAGGGTCTCAGC
 TGTGAAAGAAGATAGTATCAGCTAATTCTTGATGTGCAGTGAGGAGGCA
 GTGAGATAGTGCAGGTAAACTATAAAACAATTGTCAATGAAACGCATCA
 CAGTGATTCTTTGGACCCACAAGCTCCAATCTTATAAAACATATCCAGTC
 ACCCACCACATAGATCATCTCACCTTGCATATCTGATTTTGTGGATCAT
 GGGGAAAAACTGCTGATTCCTAGCAAAACCCATGGCATAGGATAAGTGCA
 CAATAATTTTTTTTCTTAAATGATTTAGATGACAGTGACTCATTAAAGGG
 TTTCTTGAGGCTCTCAGAGTCGAGAGGTGGTGCCTGAAGCCACCCAA
 AGTCCCTGTCAAGGATGGCTCCCAACGCACACACCACAGGCTGCCAG
 TATGTTCCACTATCTACCCAGTAGAGCCCTGCCAGTACGTTCCACTGTC
 CCTTCCCTAGAAGAGGTGACTGTTGTTCAAGTCCAGAAAAGCGGGCTC
 CCCAAAACAATGCAAGGACCCACCTCTCTCTGAACCTCACCCACCCTAGT
 TTTCTTTAAAAATCAATTTACAAGAAGATCATGTGAAGGAAAAGGTTGG
 GTGATATTCTAACCCAGTTAGCTGTTTCTCAACCAAGTTCTCTTTGAAA
 AATTCAACAACCACCTTTGGGGAATTATTTACAACAGAGGAGTGAGGATG
 GGACCAAGGATAGTATTGCCTATGTTGGTGGAACAGGGTTTTTTTCTG
 GATTACCAAGAGATGGTATGCATTGCTCCAGAAGCTAAATATCTTCAG
 GCTTCAATGGTGGCCTTCACCTGAAAATGTTATCCCTGTTGAAGCTTTC
 AAGCCAGTATTTTATAAGAACTATATTTTCTTGGTGAAGTGAAGCATT
 ATAATGATGACTATACAGGTTCTTGAGTGAAGCCATCATTAGCATT
 GTCAATTATTTTGTGTTAGTTGCATCTCCATAGCAGCTCACATTCACAATG
 TGCTTTGCAATTGTTCCCTTAGCAATAGCCCTCAAGATTCTCAGGAGGA
 GAGGTTAATCCGATTAAACATTTCTGTGAAGCCTAGCGAGATTAATCGC

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AAGAGTTTTAAAAATTAAGTAAGGACGCCGGGAAACAAATCAATCCCAGCA
 AACATTTTGTGGGATTTATCATTCAAGCAATTTTACAGTTATCCCTGTC
 AAATACATTAAGTGTCAAAATTTGGGCATAGGGGGAACAAAATAATAAAC
 CCAGCCAAAACAGAATAATCCCTGTTTGTTCATGTTGGATAAAAAAGAC
 ATTACTATTGGTGTAAAGGAAATTAGATACATCTTCCATTATTTAGTAAAA
 TTACCATAACTCTAACTTTGTGGCTTTAGGCAGTCTAGTCCACAGGCAG
 GAAGGAGGTTTGTTTTGGCAAATGACTGTTATCATCTTCTGTTTCAAAGC
 TAAACCATAAACTAAGTTCCTCCCAAAGTTAATTCAGCATATGCCAGGA
 ATGAACAAGGACAGCCTGGACGTTAGAAGCAAAATGGAGTCAGGTAGGTC
 AGATCTTCTTCACTGTCTCAGTGATGGCAGTTTCATAACTTTAAATGATG
 GCTATCACAGTTTTTCATAAAATATCTAGATAAACAGTTAAAAATAAATAA
 TTAGGTAAATGTAGTGGGATAAATATTAGTAGACAACTCACCATAATTT
 AGAATCTAAAGTTAAATTAATAATAATATTTTATTATTTGGTATTTTCC
 AAGAAAAACATATTGTAGGAAACCATTTCTTTTTAAAAAAAAGTGTCTT
 TTTAAAAAGGTGAATAATTTTGTCTAATTCAAAGTTTATTGAAAAGTTA
 TGTATAAAAACAGGTAAAAGGAACAAGGAAATAAGGGAAATGTAAAGAAA

FIG. 3 (13 of 52)

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ATTATAGAAATAAAGTGGTATTTTTTGGTAAGAAAGCTTAAAGAGAAA
ATTTTAGGTAAGAAAGAATCTTACCTAAAATTTGTGCTAGAATAAAGTG
ACTGGCTAAGAAAGGGATGTTCAAAGCTATTTATGACAAACCCACAGCCA
ATATCATACTGAATGGGCAAAAGCTGGAAACATTCCCTTTGAGAACTGGC
ACAAGACAAGGATGTCTCTCTCACCCTCTATTCAACATAGTATCGGA
AGTTCTGGCCAGGGCAATCAAGCAAGAGAAAGAAATAAAGGGTATTCAA
TAGGAAGAGAGGAAGTCAAATTTTCTCCGTTTGCAGATGCATGATTGCAT
ATTTAGAAAACCCCATCATTTTCAGCCCCAAAACCTCTTAAGCTGATAAGC
AACTTCAGCAAAGTCTCAGGATACAAAATCAATGTGCAAAAATCACAGGC
ATTCTTATACACCAATAATAGACTAACAGAGAGCCAAATCATGAGTGAAC
TCCCATTCACAATTGCTACAAAGAGAATAAAATACCTGGGAATACAACCT
ACAATGGACATGAAAGACCTTTTCAGGGTGAAGTGCACCAACCTGCTCAA
GGAAATAAGAGAGGAAACAAGCAAAATGGAAAAACATTCATGCTTATGGA
TAGGAAGAATCAATATCGTGAAAAATGGCCATACTGCCCAAGTAATTTATA
GATTCAATGCTATCCCCATCAAGCTACCATTGACTTTCTTCACAGAATTA
GAAAAAACTAATAGCCAAGACAATCCTAAGCAAAAAGAACAAAGCTGGAG
GCATTGTGCTACCTGACTTCAAACCTATACTACAAGGCTGCAGTAACCAA
ACAGCATGGTACTGGTACCAAAACAGATATATAGACCAAAGAACAGAAC
AGAGGCTCAGATATAACACCACACATCTACAACCATCTGATCTTTGACA
AACCTAACAAAAATAGCAATGGGGAAAAATAATTCCCTATTTAATAAATG
ATGTTGGGAAAACTGGTTAGCCATATGCTGAAAACTGAACTGGACCCCT
TCCTTACAACCTTATACAAAAATCACTCAAGATGGATTAAAGATTTAAAC
ATGGCTGGGCATGGTGGCTCACGCTGTATCCCAGCACTTTGGGAGGCC
GAGATGGGTGGATCATGAGGTGAGAGATGGAGACCATCCTGACTAACAC
AGTGAAACCTGTCTCTACTAAAAAATACAAAAATTAGCTGGGCATGGT
GGTGGGCGCTGTAGTCCCAGCTACTTGGGAGGCTGAGGCAGGAGAATGG
TGTGAAACCAGGAGGTGGAGCTTGCAGGGAGTGGAGATCACGCCACTGCA
CTCCAGCTGGGCAACAGAGTAAGACTCCATCTCAAAAAAAAAAAAAA
AAAAAAGAGGATTTAAACATAAGACCTAAACCATAAAAAACCATAGAA
GAAACCTAGGCAATACCATTGAGGACATAGGCATGAGCAAAGACTTCAT
GATTAGAACACCAAAGCAATTGCAACAAAGCCAATTGACAAATGGGAT
CTAATTAACTGAAGAGCTTCTGCACAGCAAAAGAACTATTGTGAGAGT
GAACAGGCAACCTACAGAATAGGAGAAAAATTTTTCAATCTATCCATCTG
ACAAAGGGCTAATATCCAGAATCTACAAGGAATTTAAACAAATTTGCAAG
AAAAAAAAACCCATCAAAGTGGGCAAAAGATATGAACAGACACATCTC
AGAAGAAGACATTTATGTGGCCAAACATGAAAAAAGCTCATCATCA
CTGGTCATTAGAGAAATGCAATTTGAAACCACAATGAGATACCATCTCAT
GCCAGTTAGAATGGCGATTATTAAGAGTCAAGAAACAACAGATGCTGGA
GAGGATGTGGAGAAATAGGAATGCTTTTACACTGTTGGTGGGAGTGTGAG
TTAGTTCAACCATTTGTGGAAGACAGTGTGGCAATTCCTCAAGGATCTGGA
ACCAGAAATACCATTTGACCCAGCAATCCATTACTGGGTATATACCTAA
AGGATTAGAAATCATTCTATTGTAAAGACACATGCACATGTATGTTTATT
GCAGCACTATTCACAATAGCAAAGACTTGGGAACAACCTAATGCCCCACC
AATGATAGACTGTGTAAAAAATGTGGACGTATACCCCATGGAATACTAT
GCAGCCATAAAAAAGAATGAGTTCATTCTTTTGCACGGAACCTGGATGAAG
CTGGAAGCCATCATTCTCAGCAAACCTAACACAGGAACAGAAAAACCAACA
CTGCATGTTCTCACTCATAAGTGGGAGTTGAACAATGAGAACACATGGAC
ACAGGGAGGGGAATGTACACACCAGGGCTGTGAGGAGGTGGGGGGCAA
GGGGAGGGATAACATTAGGAAAAATACCTAATATAGATGACGGGTAAATG
GGTGCAGCAAAACCACCATGGCACATGTACACCTACGTAATAAACCTCCAT
GTTCTTCACATGTATCCCAGAACGTAAAGTAAATTTAAAAAAGAAAGAA
AGAAAGAAAGGATGTTACAGCAAAACCAGAAAGTCCAAGCATGTCTATGA
ATAGTCTGTGTAAGTCAATAAGAGGATTTATTTAAAAAACTTTTATA
TGATAAAGTTGTCTATAATTAAAGGGAAATTTAATGGTCTTTCTAGAGA
TTGGGTTGATGTTAAAAAACTACTTATATATTAAAAAATTGGTTAGAACA
ATGAAATTTTCTTACGGGGTTGATTCACTCTTAATAAATTATAAGAGACT
TAAGAATTTTTTTTAAACCCAAAGTTGAGCTTTTATTGCATCTTGCTGTT
TTAGGTTTTCTCTCCCTTTTAAAGGGTGGGAAATAGTAATGCCCTCCTT
CAACTCCCTTCAGCTCATATACGTTTTTTTACCCTCAGATTCTGTTTGTG
TGCTCTGATGCTAACAATGTTTTCTTAAAGGTCTAAAGGAAATGTTTTCT

FIG. 3 (14 of 52)

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TCCAACATAATATTCTGTCATTGCGAGAAGGTCTTTCTTTTGCCTTTT
 GTAACCTGGCTTAACAGATTTTATGTTTTATTGAAATAATTTCTATGCCAT
 TATTATTAAGTTTTGGTTTGGCTTAGAAAACACTGAGATTAATACAATTTT
 TAAAAATTATGATTATTACATCCATATATCTTTATGTATGTGCTTTTAA
 AGTCCTTGTGACATTGAGTTCTAGGGCTTGACTCCTGGGTCTTAAAAGGA
 CAAGTCCTGCTAAATCTTAAATACTGACAGCAATTAAAGGCTCATCTTCA
 GGACTGGTAGAAAATGCCAATCAAAATAAACTGCATTCTTGAAACACAGA
 GCCAGAAATTAAAGCTATTCAACTCAAGGCCAGGAAGTATAGTGGAAGA
 GGTGGGTGTGTGAGATTGTAAGGGCCAATTTTGAGAGATAAAATAAGTTC
 AATTTCTCTATAAATTAATCATAATCATTGATGTCCAAGCCACACTGATG
 CAAGATCAGCATATGGGTCTGTGTGAGATTAAACAGGTTTTCTTGAAGC
 ATTAACCTACTCCTTAATAAAGGTTATAGAGGTTATAAAGGCTTCTGGA
 AGTTATAGTATGGTCAAGATAAAAATTTCATAGATTGTTAATACAATTT
 TGGAAAACAAATTTAATTGGCTTCTTGCTGTTTTTATTAGGGCTTATTGT
 TTGAAAATTAAGTCTCGTCTCTCAAAGAAATGAAGGCTTTCACCTTTTTT
 TTTTTTTTTTTAATCCTTGAGTTATCACTTTGGTCAAATGAATGACTTA
 TTTTACAATGACCTTTTATCAAGTGTTTTTAAACCTTTCAAATTTGACAAA
 CTTTCCAAAATCAAACCTACAAATTAATGTCTTTTTATGACCTAATGAATCC
 TTTAAAATACTAGGTTCCCTAAAGTCCAAAAAATAACATAA
 TGTGGCTTATTTGGTATAAAAAATTTACAAGAAACATTGTCAAATATAAA
 ATATTGTGTGGTTTTGTTTGGGCTGTATTTGTATAAATATGTTATTGGTA
 TGTGTTCCAAATTAAGGAACTCCTATAATTCTGATATGACTTGGTGT
 ACATTATCAGTAATAATTATAATTGTTATGGTAAATTAATTGTGTGCCATG
 GAGGTAACAAATTTCTCATCAAGTGTGTCTTTGACTATGGTTGCCCTAA
 AACTTTTGGCATTACAGACAATTGTCTGCTTTGGTCTCTTTAGAAG
 GTGGTTTTATAATCAGCTATAAACTCTAACGGGTGCTCTTGAATGCAGG
 CTTAAGATAGCTTTGGAGACTGTGACATCAGAATAGAGGAAAACTTTCA
 GTATTCTAGGAGTGTGAAATATTATGAATATCAAGCAAAACAGGAATT
 AACTTCATAGATGGAACCTAAAAGAAATGCTGAAGTAATCTTTTTGACTTTT
 TTTCTTAGAATGTTGATCCTTCGTTTTGTTTTTCAGAGTCNAGGAAATTT
 TTCTGTTGAGATATTGACAGCTTTAACAATTAAGTATACTCCAGTGAACA
 CAATTTGGAGCA

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ATCTAGTCATTCCCCAGCCTGACCAATTCAATGGCCCCCATCTTAGTTAA
 AATTCCTCACCCTGACAAGGCCCATCTACGCCCTGACCTCATGCCCTC
 CACTCTCAGATTTGCACTCACCCTGCCCACTCAAGGGCTTCCCCAGGTT
 CTTCTTAGATTCCACCGATAGCTCAGGGACTTTGCACATGCTACGGTCT
 CTGCCCTGGCTCCTCCCAGATCTTCTCATGCCCTAGCTGCTTCTCATCAGC
 ACCCTCAGAGACTGTCCCTGCCCCACCTCTCCAGGTTCCATACCTGCCA
 CCTCCCCCAATCACGTAACAGTTTCTTACAGAGCGAGTTACCATCCCA
 GTATTTCCCTAACTTATTTTTGTGACTGGTCTGTGCCCTGTCTCCACCA
 CAAGAACATAAGCTGCATGTGAACAGGAGCCTTGTCTATCTTGTACCCCC
 AGTGGCTGTGACATAACCTGATACACATTAGATGCTCAATGATGTTTGAT
 GAATGAAGTGCTGGTAGTCCAACTGTGTTTCTTGTCTGTGTAAGTATGT
 CTGTTGTGGTTTTCTAAGAACCTACAGCTCTCCACTGTGACTCCTGTTT
 TATGGTCTGATTTGCTGGACTAGAATCCTAACCTACATGCTTACTCTTA
 GTGTCTCTCCCCAGAGGCTGAATCCAGTCCCTAAACCTCCACCAAATGG
 CTAAGACCTAGCTTCCAACAGACAGGCTACGCTGAGACCTCAGCACCG
 CCTTCTGCGGTCTCATCTTAACGCATCCTTCAGGGCCCAGCTTAAATG
 TCTCTTCTCAAGGAAGGCTATCCTCTTTCTGCCCTCAGTGCTCTCCAT
 GCCTCCTCTATGCCTCCATGCCTGCTTTCAACCCTGCAGAAAGTGGAGAAA
 TTGCTAATCTGCTGTGTTGACACTGTGCTGGGGTGCTTGGGCCAGGGAG
 CAGGCTGGTGGTGTGCTGATAGCCCGTGGCTGTGCCAGGTCCATGCTCA
 CTTCTGAGCCCCAGTGGAGTAGGCTCCCTTTCCCTTATTGCAGCACTCA
 GAGGAAGGACGTGCTTCTTAGGACAGATCTGGCCAACCTCTCCCTCGTGA
 GAGAAGGCCAGCCATCCTCTTGCCCTCTTTCTTCTCCTGCCCCCGAGT
 AATAAAGGTGCCTGGTCAGAGCCTTCTAGAAGGAGACCCAAACATCCACC
 ACACATTCCCAGTTCCAACCGTCATCCACATGGCTGGCTGTGCAGGTAAA
 CGCAGAGTCTGTTTCAACACACCCCAACCATCTAGTATTGGATGGGAGGACA
 GTAGCGTGACACTCTTCTCCAGCCTTGAGCCCTACTGTGGGGCCCCACCCA

FIG. 3 (15 of 52)

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ACCCAGATACCAGAGGAGCCCTGTACTGGGATGCTATTGGATGCTTGGTCC
AGTCATGTACAAAGTTAGCCCTTTGTTATATAGAGTTAGCTACGTACATC
TTCCTCTGTAGGGAACCCAGAGGGGAGAGAGATATGTAGTAGGATTTA
ACCTGCAAATCCTCTGCTGAGCACCTGCACTACATACAGTGGGTAGCAT
GTGGTAGGTGCTCAATAACTATTGACCGATAGATTGAATACAGGTAGGAT
GGTGACACAATCTAAGATCCAGGGGTGGGGAGACCACACGCTTGGTTAG
GGAGACCCAAAGTGGACCGTGTGGCCAGAAGAGTCCCGCACTGCACTCTA
GTGACAGTGCAGAAAGTCACTGTGGGAAATCTAGAAGTTTCTACAGGTTG
CTATTTTCATCATAGCACTGTGCGAGGCCAACCCCTTCTGCTCCACTGGCTG
TTGGGAAAAGCTTTCTCTTTTCTTCTCTAGCCAGGGAGCTCTCAAAGTGT
CCACTCTCTCACCTCCACCCAGGCGTCCAGGTGTGGAGGACACTTGCCGG
CTGCTTGTCTGCTGACTCATCCCTTGGTTTCACTTGGAAAACCTACCACC
AGCTGGCCTCTTTCCAAGCATCAGCCTCCTCATTTTCTTAATCCCTTAGG
TGTGATCTCACCTCCACACAGTAGATTGCCTCAAGGCCAATTCCAATAT
GAATAAAAAATGATTATTTTGTCTCTTCCAATCTTCTTTTAAAAATATTA
TTTTATAATTTCCCTTTTAGGAGGATCACCTAAGTGAAGACTATTTTACCT
AAGAAAATGTTAAAAATGTAAAGACATGGTTGTAATCTGGGGATTCTGTTA
AAATGGCTAGCAGACAGAAGTCAGACGACAGGCTAGAAATGTGTGAAGAG
TGGTTGCCCTTTGAAAGGCGGAGTTGGTAATGATTTTCTTCCATTTTCCA
TGCTTTCCAATTCTCTACAAAGGCCTTAATATTACTTCGATAACCAGGAC
CTCTGATAACCTGCCCCCAGGAGTAAAGACTTAGCTGGGAAAGTCAGCT
TCATGTGAGGTAAAGGAACCAGGTAATACACAATTCCCAGTGCCAAGT
TCGGGTGTGAGGCTGAGCTTCTGCTGTGGGAGGAAAGAGAAAGAAG
AGAGAACTCCAAGATCCAAGAGATCCAGCAAGAAGGCTGGAGTCTGAGG
ACGCAGAAAGCTGAATGGCACAGTTACCACTATTGTGCTGAGGTTCTGTG
GCCTCTGGGTCTCTTGACAACCTGGGCAAAGACCCACAGAAAATCTCTCT
AGACCCTACCTGTGGGAGGGGAAAGTGCTTAAGATCATTACAGGACAGC
CACCTGGACCTCAAATGGCTTACAGTTCCTTCATCCAGAGGGTCTTCATT
TAGTACATACCAGGTGCTAAGCTGGGTGCTGGAGACATGACGGGGAACCC
ATTTACCATGGCTTTGTTACTGTGACATTACATCTAGGGAAGGCCAGCA
AAGGGGAGGGATCGAGGAGAGCTTGTAGGCAGAGAAAATACCCAAGGGC
AAGGGAGAAGCCAGCCTGTTCTGAGCACACACAGTGGTTCCATCTAACTG
GGCCTCAGTGCCAGGTTGGACTGGAGATGGGGCTGAGGAGCTGTACAGA
GCATTCTGGACACAGATGTCACATAGTCCCTTGAGGTTAGGGTCTTAGG
CATGGCAGCATTGCTTTGAGTTTTTCTTTTGTAAATGTTGCCATTCTGA
CAATGTGGAAGATGGGTCTTGACAGAGAAGGGCAGGGCTGTGAGACCAGT
TAGGAGACTAAGATGTGAGCCAAGGAAAATGAGGAACACCTGAACACTGG
GGCAGGTGCAGGGCCAGAGAGAAGCAGATGGCTTCTGAGGTTTTAAGT
AGGTAGAATCAAGGCAGCTGGTACAGATCTTTTATTACATATAAACTGGA
ATAAGCCATCTGTTCCAAGACAAAAGAGTAGGCGGAAAACAATACAAGAC
AGAAAATGGAATTAGAACAAACCTGGGAGGAATGTGGAATTAGAGTAGAGA
GTCCAACACTGGCTGCAATCATAAAAAATGTAAACAAACAAAAATTTGCT
AGGTGTGCTTACTTAGAAATAATTAGCTGTCTATTAAGTTCACTTGTGT
TATGGCTTAAATGTGTCCCCCAAATGTGATGTGTTGGAACTTGATCCC
CAATGCAACAGAGTTGAGAGATGGGACCTTTAAAGGTGATTAGGTCATA
AGGGTTCTGCCCTCATAAATGAATTAATACTGTATCATGAGAGTAGATT
CCTGATAAAAGGATGATCTCTGCCTCCTCCCCACAGCCCTCTGTGCATG
CTTTCCTGCCTTTCCACCTTCTGCTATGGGATGACACAGCAAGAAGGCCC
TCACCAAATGCAGCTCCTTGATCTTGACTTTCCAGCCTCCAAAACCTGTA
AGCCAAACAAATTTCTGTTTATTATAAATTACCCAGTCTCAGGTATTCTG
TTCTAGAAACACAAAATGGACTAAGATCATTAAATTATCATTTTTTATCA
GACTGTTGA

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AAAATATAACAGAGAGTAAGAGGAAAATTACCTTCTTTCTTTTCTTTTCT
CCTGCCTGACCTTATTCACCTCCCATCCAGAGCATCCATTTATTCCATT
GATCTTTACTGACATCTATTATCTGACCTACACAATACTAGACATTAGGA
CAATGTGGCCTGCCTCCAAGAACTCAAATAAGCCAACCTGAGATCAGAGA
GGATTAATCACCTGCCAATGGGCACAAAGCAACAAGCTGGGAGCCAAGTC
CCAAAATGGGGCCTGCTGCTTCCAGTTCCTCTCTCTGCAATTGATGTCA
GCATTATCCTTCGTCCAGTCTGTCTCCACTACCACTTTCCCCCTCAA

FIG. 3 (16 of 52)

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CACACACACACACAACAGCCTTAGATGTTTTCTCCACTGATAAGTAGGTG
ACTCAATTTGTAAGTATATAATCCAAGACCTTCTATTCCCAAGTAGAATT
TATGTGCCTGCCTGTGCTTTTCTACCTGGATCAAGTGATGTCTACAGAGT
AGGGCAGTAGCTTCATTGATGAACCTATTCAACAAGCATTATTCACTGAG
AGCCTTGTATTTTTTCAGGCATAGTGCCAACAGCAGTGTGGACAGTGGTGC
ATCAAAGCCTCTAGTCTCATAGAACCTTAGTCTTCTGGAGGATATGGAAAA
CAGACAACCCAAACAACCAACAAAAGAGCAAGATGCTGCAAAAAAAAAAAAA
AAATGAATAGGGGTGCTAAGATAGAGAAAAGTGGGAGAGTGTCTATTTAGAC
AAAGTGGTAAAAACAAGCCCTTGTGAGATGAGAGCTGCCGACAGGAGG
GGGCGGGTCATGGTTGTGGGTTTTTGGGTAGGACATTAGAGGAGGGGGC
GGGTCTGTGGTTGTGGGTTTTTGGGTAGGACATTAGAGGAGGGGGCGGGT
CGTGGTTGTGGGTTTTTGGGTAGGACATTAGAGGAGGGGGCGGGTCTGTG
GTTGTGGGTTTTTGGGACATTCAAAAGAGTCTGAATGCACCCAGGCCTAC
AACTTCAAGATGGTAAAGGACAGCTCCAAGGATCAGAAGAAGCATGCTTG
GAACTGGGGCATTGTGAGAAGGAGGAAAAATATGCAGAGACTAGTGTCTTG
CAGAGCTTGCATGTGGATTTTCAATTTGAGGTACAATGAAAACCCATTAATG
GGTTTTCACACAGTGCATGGCCTGACCTCACTTATATTTCTAAAATAGA
AAACAGATCAGAAGGAAGGCAATAGAGAAGCAGAAAGTCCAATGAGGAGG
TTTACAGCAGTCTATGGGGGTGGGTAAGGAAAAGAAGTGAAAGAAACA
GACAGAATTGGGTTATATTTTGGAGATAGAACCAACAGAAGGAAGAGGAG
AAACAACATTTACTGAGAAGGGAAAAAGTAGGAGAGGAATAGGTTTGGGA
AATAAATCCTGCTGACATTGGAAACCCCAAGGAAGCCTCAAAAGTATATT
TACTTGCCTTTAGATTTAAAAGAATAGGAAAGAAGCATCTCAACTTGGAAAT
TTGAAATCTATTTTCCATAAAAAGTATTGTTAAATTCTACTCATACTCAC
AAGAAAAGTACATTTCTAAAGAGTATATTGAAAGAGTTTACTGATATACTT
AGGAATTTTGTGTGTATGTGTGTGTGTGTATGCGTGTGTGTGTGTTAAC
CTTCAATTGTTGACTTAAATACTGAGATAAATGTCTATAATGCTAAAT
TGATTTCCCAAAGGTATGATTTGTTCACTTGGAGATCAAAATGTTTAGGG
GGCTTAGAATCACTGTAGTGCTCAGATTTGATGCAAAATGTCTTAGGCCT
ATGTTGAAGGCAGGACAGAAACAATGTTTCCCTCCTACCTGCCTGGATAC
AGTAAGATACTAGTGTCACTGACAATCTTCATAACTAATTTAGATCTCTC
TCCAATCAACTAAGGAAATCAACTCTTATTAATAGACTGGGCCACACATC
TACTAGGCATGTAATAAATGCTTGCTGAATGAACAAATGAATGAAGAGCC
TATAGCATCATGTTACAGCCATAGTCTTAAAGTGCTGTTTCTCATGAAGG
CCAAATGCTAAGGGATTGAGCTTCAGTCTTTTTCTAACATCTTGTTCTC
TAACAGAATCTCTTCTTTTCTTCATAGGAGATGCCTGAGATACCCAAAA
CCATCACAGGTAGTGAGACCAACCTCCTCTTCTTCTGGGAAACTCACGGC
ACTAAGAACTATTTCACATCAGTTGCCATCCAACTTGTTTATTGCCAC
AAAGCAAGACTACTGGGTGTGCTTGGCAGGGGGGCCACCCTCTATCACTG
ACTTTCAAGATACTGGAAACCCAGGCGTAGGTCTGGAGTCTCACTTGTCTC
ACTTGTGCAGTGTGACAGTTTATATGTACCATGTACATGAAGAAGCTAA
ATCCTTTACTGTTAGTCAATTTGCTGAGCATGTANTGAGCCTTGTAATTCT
AAATGAATGTTTACACTCTTTGTAAGAGTGGAACCAACTAACATATAA
TGTTGTTATTTAAAGAACCCCTATATTTTGCATAGTACCAATCATTTTA
ATTATTATTCTTCATAACAATTTTAGGAGGACCAGAGCTACTGACTATGG
CTACCAAAAAGACTCTACCCATATTACAGATGGGCAAATTAAGGCATAAG
AAAATAAGAAATATGCACAATAGCAGTTGAAACAAGAAGCCACAGACCT
AGGATTTTATGATTTTCAATTTCAACTGTTTGCCTTCTACTTTTAAGTTGCT
GATGAACCTCTTAATCAAATAGCATAAGTTTCTGGGACCTCAGTTTTATCA
TTTTCAAAATGGAGGGAATAATACCTAAGCCTTCTGCCGCAACAGTTTT
TTATGCTAATCAGGGAGGTCAATTTGGTAAAATACTTCTTGAAGCCGAGC
CTCAAGATGAAGGCAAAGCACGAAATGTTATTTTTTAATTATTATTATA
TATGTATTTATAAATATAATTAAGATAATTATAATATACTATATTTATGG
GAACCCCTTCTCCTCTGAGTGTGACCAGGCATCTCCACAATAGCAGAC
AGTGTCTTCTGGGATAAGTAAGTTTGATTTTATTAATACAGGGCATTG
GTCCAAGTTGTGCTTATCCCATAGCCAGGAACTCTGCATTCTAGTACTT
GGGAGACCTGTAATCATATAATAAATGTACATTAATTACCTTGAGCCAGT
AATTGGTCCGATCTTTGACTCTTTTGCCATTAACTTACCTGGGCATTCT
TGTTTCATTCATTCACCTGCAATCAAGTCTTCAAGCTAAAATTAGAT
GAACCTCACTTTGACAACCATGAGACCACTGTTATCAAACTTTCTTTTC

FIG. 3 (17 of 52)

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TGGAATGTAATCAATG1. TCTTCTAGGTTCTAAAAATTGTGATCAGACCA
TAATGTTACATTATTATCAACAATAGTGATTGATAGAGTGTTATCAGTCA
TAACTAAATAAAGCTTGCAACAAAATTCTCTGACACATAGTTATTCATTG
CCTTAATCATTATTTACTGCATGGTAATTAGGGACAAATGGTAAATGTT
TACATAAAATAATTGTATTTAGTGTTACTTTATAAAATCAAACCAAGATTT
TATATTTTTCTCCTCTTTGTTAGCTGCCAGTATGCATAAATGGCATT
AGAATGATAATATTTCCGGGTTCACTTAAAGCTCACATTACACATACACA
AAACATGTGTTCCCATCTTTATACAACTCACACATACAGAGCTACATTA
AAAACAATAATAGGCCAGGCACGGTGGCTCAGACCTGTAATCCCAGCAC
TTTGGGAGGCCAAGGTGGGAAGATCACTTGAGGTGAGGAGTTCAAGACCA
GCCTAGGCCAATAGTGAGATCTCATCTCTACAAAAAATGAAAAAT
TAAAAATGAGCTGGACATGGTAGTACACACCTGTAGTCCCAGCTACTCG
GGAGGCTTGAGGTGAGGAGGATCACTTGAGCCTGGGAGATGGAGGCTGCAG
TGAGCCATAATCACACCATTGCACCCCAACCTGGGCAACAGAGTGAGACC
CAGTCTCAAAGATAAATTTTTAAAAATGTTAAAAATATATAAAAGAGA
ATTTTAAAGAACAATAATAGATCAAAGCATGGATGCAAGATATATTTA
GTTGGAAAATCAAGGTTAAAAATCAAGGGATCTTGAATTAGGTGTGGTAG
ATTTGGGTAAGGAGTAGTCTAAGATGACCCTGTTTCTTGGTACTGGAGAC
TGGATGAGTGGCAGCGTCTTAACCATATTTTGGTAGAAATATGGAGGTC
TTCTCCATTCCAGGATGAATGATGAGTAAAATTTAGGCATGTAATTTGA
GCTACTAGAAGGCACTCAATTGCAGATGTACAATGGGGAGATGATAACC
TATCTGGAATCAGAAAAATAACTGTATATAGATATGAAAGACATCAGTA
GGTATGTAGTAGATAAAATCCTAAAAGTGATGTCAAAGGGAGAAGAGAAG
TATATGGTGAACACTGTTGTTTGTCCATGCAATTGCCATCTCTTCTCTT
CCTTACTGACAGAACCCTGATTTCACTGAGAAGTCAACATGCCCTTCCCC
AATTGATGAATCCAATTGGTTGAAGATTATGTTCAATCTATTCTTACATG
ACTAAGTCACGTTGACTTAATCCTATCAAATGAGATGTCGATCTGGAAAC
AACTTCTGGAAAAGATTTTCTACCTTGATAAAATAAAGAGCCATATAGAT
GGTCCCTTATCTTCTTCTTCTTGAATGAGATATGTTCTATGAGGAAGT
GAAGCTTAGAATCTGGTTCAGCAACTTGCAACGACTGGGAAGTCAGAGCC
ACACAATGAAGAATGCAGAGTGGGAAGGAGAAAAAGAGCCAGCATCTCTGA
CAACATTGTTACACCGAGAACCCTACCTCCAGATTTTAAAGAAAACAAGAAA
TGCTACTGTTATTAAGCCATTTCACTGGGTTTGCTATGACTTGCAGTCAA
ATCTAGCTTAACTGATACAGAGCACCACAGAGAACTGGTCTCTCATTGT
CTCATCTGTCTTTCTAGCAGCCACGACTTTCTAGGGTTTCTTAGCC
CAAGTCTGGCTAGAGCAAGACTAAGTAAGACTTGATTCTTAAATGTCCTT
TTGTTTTAAGAAATATTAAGAATTATTTTTATATTAATATTTTTAAGA
AATAAGGAAATACAAACACTGAGCAAGCAACACAAATTCAAGAAATCTT
AAAAAGTATAATAGCTGCTCAGTCTCTGATTAAACAGTGAAATATGGAATC
ATTGTAGAAATGGCCTTGGAGCGTTATTCTCCAGGCCAGCTATCCTTAT
GGTCTGCCCCACCTCCCTCATTGCCTAAACAGTAAGAGAGTCCCATGGTG
AGACTCAACAGTCTTAGCACAGAACTTGTTACAGTCTATTTCTTTCTTA
CAGTCTTATATATCAATTCCAAATCAATGAGAGTAAAGCCCAATCCCTGC
CTTTAAACCCCAAGGACAGAAGCCCAAAGCCCAAAGATATCCCTAACCT
TCTCCCCCT
>Contig28
CCTGTGCTCCCTATGTTTAAAGCTGGGGATCTCTTTTCTGTGTCTAA
TTATTTTCTCATTGGCTTGAAAAATCTGATAAAACATTTTAGGACTGTG
TATAAAATAGAATTAGCCAAGTGCAATGTCTTTATTCAGAAGAAATTTCA
TGGACGTTGTGCCTACTCTCTGGCTTCTGGCTTCATGGCTTCCAGAT
CCCACAGTAAGCTCTGGATAGTAGAAGTTATAGTAAGACTGACTTCTAAA
TAAATGAAGTGACTTTAACCTTACTGATATGGCTTAAAGAAAAGGAGTGG
CCTTTAAGATCCATGAACCTTCTCAAACAAAAGTGATAACGTTATCTCCAT
GCATATATAATACTAAATATAATGCAACTGAGAGAAGTAGGCTGTGGTAA
GAAAGGAGACCCAAAGTGCCATCTGAAGGCAGCACTTACCACCTCTGCTTCA
TCCCACCGAGGAAACAAAGCATGAGTATTGCCAGATTTTCTTCTGTTTCA
AGAAAAGCCAGAAATCCAGGTTTTTGGCGTGAATGTCTGATTTTAAATGT
TGGGAAGTAAATTTATATTTTGAATAACATTGTGTGGGACAAGTGAACCTT
GTATGTGGAACCTGCTTTCTCCAGTGGCGACAGTTTGGACCGTTGATAC
TCAGCAAGTTTCAGCCAAGTGCGCCTTGTCAATGTCAATCAAGGTGAT

FIG. 3 (18 of 52)

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GTGTGATTGGTCAAACAATTAGTTTTGCTCAGCATCTCGTGTGTTTTCA
AGGACCTGAGGGTTCAATTTGCCCATGCAGATCTTGTAGTCCTGTTTATTC
TATTAATTTATCTTGCAAATCTATAATGTTTTATTTAAGCAGCGAGAGC
CGTGGCAGCCTTTGGTCTGGACCCTTTCTAATGATCATTTAGTATCAGGC
TATGTGGGAGTTGATTGTTTTGCATTGCCTGAAAGCCAACAGTATCACTC
CTCCTCTAGGTGTGGCAGAGATGTGAGAGAGGGAGACTGACAGTCTGTGG
GTGTGTATGCAGTGTGTTGGGGGAAGCGAGGCACAGGGGACAATACTGTGGT
GTATAAACTAGTCTAAGGTAGCATCAGGAAGTTCATGAAGCCAAAATGA
TTTTTCATAACAGCACAAGACATTATTTGTTTTTGCCTCCCTCTCATTTTT
TTTTTTTTTTGAGACAGAGTCTTGCTCTGTCTCATCCATGCTCGTGTGCAGT
GGTGCATCTCGGCTCACTGCAACCTCCACCTCCAGGGTTCAAGCAATTC
TCATGCCTCAGCCTCCTGAGTAGCTGATTACAGGTCTGCACCACCCCGCC
GGCTAGTTTTTGTATTTTTAGTTAGAGATGGGGTTTTGTAATGTTGGCCAG
GCTGCCCTGTCTATTTTTTTTTTACTAGTGTCCAGTGGAGTTTTTTAGGGG
CTACATAACATGATACTGTCTAATTAATCTAATGGCTAATGAAAGGGATATG
TATATGTTTTTGTGTTTTAAACAAACTTCTTTGGGGTCCCTCAATAATTTT
TAAGAGTATAAAGGGGTCTGAGATCAAAGAGTTTGAGTCTGCTGGACT
GGGACAGTGGTTGTCAACCCAGATTGTACATTAGGGTCTCTGGGAAGCT
TTAAATAGTACTGATGCCCAACCTTACCGCAAACCAATTAAGCCAGAAT
CTCTGTGGATGAGAAGTCTTCATTGTCTCATCATCACCATGACCATCATCAT
TGTCACCGTCACTACACCATATCATCATCATCATATCATCTTCATTATC
ATTGTTAGTATCTCCATCACCATCATCAGCATCACCATTATTATCATCAT
CATCATCCCCACCATCATCCTCATCGGAACCTTCACCTGCATGGAGGACAA
TCCACTATGCATTAGGTGCTATGCTATTTGCTATACTCCTTATTCTCACA
ACTGCCAGAGAGGCTGATATTATCTCACTTTATAACAGGAGGAATCTGG
ATCGGAAAAGTTAAGGTAAGCTAATTCACAGAGCGAGAAGAGATAGAGCC
AGGATTCGAAACCAGTTCTCTGCTACATCAATGTTCCAGTCCCTTGCACT
ATTGAGAACCCTCTTTAGTTATGCTTTCAACCCCTCAACACCACAGTAAAT
TTTTCTTTTTTAAAAAAAATTATACCTTTAAGTTATAGGGTATATGTGCA
TAATGTGCAGGTTTGTACATATGTATACATGTGCCATGTTGGTGTGCTG
CACTCATTAACCTCGTCACTTACATTAGGTATATCTTCTAATGCTATCCCT
CCCCGCTCTCCCCACCCCATGACAGGCCCTGGTGTGTGATGTTCCCCACC
CTGTGTCCAAGTGTCTCATGTTTCAGTTCACCTATGAGTGAGAACAT
GTGGTGTGTTGGTTTTCTGTCTTGTGATAGTTTGCTCAGAATGATGGTTT
CCAGCTTCATCCAGTCCCTACAAAGGATATGAACCTCATCCTTTTTTATG
GCTGCATAGTATTCCATGGTGTATGTGTGCCACATTTTCTTAATCCAGTC
TATCATTGCTGGACATTTGGGTGGTTCCAAGTCTTTGCTATTGTGAATA
GTGCCACAGTGAACATTCATGTGCATGTGTCTTTATAGCAGCATGATTTA
TAATCCTTTGGGTATATACCCAGTAATGGGATGGCTGGGTCAAATGGTAT
TTCTAGTTCTAGATCCTTGAGGAATTGCCACACTGTCTACCACAATGGTT
GAATTAGTTTATAGCCCCACCAACAGTGTAAAAGCATTCCTATTTCTCCA
CATCCTCTCCAGCACCTGTGTTTTCGTGACTTTTTAGTGATTGCCATTCT
AACTGGCACCACAGTAAATTTTTATAGATTTTATAAGCAAATTGTATTTA
CTGTGCAAGAATTGGTTTTATTTTTTAAACCATGTGTGCAAACATACAAT
GGTTAATTGTGATATTTGCTCAGTACAAGATCATCAGATCACTACACAGA
CTTGAGGTAATTCACCTAAAAGCAAAGAGAACTGACCCACATTAACTG
AGAAGTCTTTACTTATTTATTCCTATAAACGAGCCAATATGAAGAGAAG
GCCTTAATGTGGTTAACTATGTAATTTTTTTCTGACTTTTTGAAATACTG
AGAAGAGCTCATGACTCTCCCATCTCCTAATTCTACCTGGTGGATTTTA
GACTGACCACAACCTCATGGGTAAATGAGGGAAGACGAATAAGAAACCTTG
CTTTTTTTCTCCTTGTTTTTGGCTGGCTGCAGTGGCTCACACCTGTAA
TCTCATCACTTTGGGAGGCCAAGGTGGGAAGATCACTTGAGCTCAGGATT
TCAAACCTGGCCTGGGCAACATAGTGAGACCCCATCTCTAAAAA
AAAAAAAAAAAAAGGCGACAGGCGGTGCGTGCCTGTAATCCTACCTACTC
AAGAAGCCGAGGTGGAAAGATCACTTGAGCATGGGAGGTCAAAGCTGCAG
TGAACCTTGATTGCACCACTTCATTCCAGCCTGGGTGACAAAGCAGGACG
CTGCCTCAAGAAAACAAAAACAAACCTTAATTTTTTGGCTATTCTTTTC
TGGTAAGAATGGTATAGAGATGGGGATGAGGATGGCTATTGTATGAGAGA
GCAAACAGGGTCCAAGCAGTGCTCTGGGCTGTCTAAGGACCAGTAGTCAG
CTTAACCTCTCAAATTTCCAGGGAAGGAGTTTCGGAGTGGTAGAATATCCT

FIG. 3 (19 of 52)

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GGGTATGCCCAAAGCATCACCTTGCAAATAGCCTGTCTATGAATAATTTG
TTCATTTTGTATGACTGGAACTGGCTTTGTGTATGCCAGAGAATGGGGG
CAGGAAAGAGAGATTGGTGTCTTGAGCTCTCTGTGCCTCTGGGGCAGTGA
TGCTTTTCTCTCATGTGGAAGGAGAGCATGACTGAAAAGGTGCACAAAT
AAGGTGTCTGTGAGAGAAATTAACCTTCCAGATACAGAGACACAACCTTC
CCCAAGAGGTCTCATTTGCTCTGCCTTTTTCCTTTTTTTTGCTTGTCT
ACCATTAATAACAGAACTGATTATGACCTCAAAGAGAGGAGAAAGCGA
CTCTCCCCACCTAGAGCTAGTTAACCACCATATCTTCTAGATATCCTT
GAGAGCAATGTAACCC
>Cont:ig29
GTGAACTCGTTTTACCTGTGTAGCAGACCAAGCCGAGACAAAATCCNTC
AGACACCAAATTAAGAAGGAAGGGCTTTATTGGGCCTGGAGCTGCGGCA
AGACTCAGCTCTCCAACACCGAGCTCCCCGAGTGTGCAATTCCTGTCCC
TTTTAAGGGCTCACAACCTCTAAGGCGGTCCACATGAGAGAGTCTGTATAG
ATTGAGCAAGCAGGGGTATGTGACTGGGGGCTGCATGCACCTGTAGTTA
GAATGGAACAGAACATGACAGGGATCTTCACAGTGTCTTTCTATGCAAA
TAACCGATTAGATCAGGGGTCTGATCTTTACCAGGCCCAGGGTGTGTCAAC
GGGCTGTCTGCTTGTGGATTTTCTTCTGCCTTTTAGTTATTACTTCTTT
CTTTGGAGGCAGAAATTGGGCATAAGACAATATGAGGGGTGGTCTCCTCT
CTTACCTGCGGGGAGTGAGCTCAAACCTCTTAAAGGAGTTACCTGCCTTC
CATCATCAGGGAAGCAGGAAATCTTGCCTTCTTGTGGAAGCAASTAAA
ACTCAAAACAAACAAAGAAAAAACAGGGAGTTGTACAGCAAAATAAACT
TTTGATTTTGAACAAATTTTGGGAGATCAGGAATCTCTGAAGGAGATGC
TTTCAGACCTCAGCAAAATTTCTCTGTTGGTTTGAAGCCATAAAGTTAGCTC
ATGCTGGTACCAACACCCAGTAGGAGATTTGTCAAAGGTAAGAGGCATCT
CCACTCAGAATCCCTTCGTGGTTACCAACATGTGAACCTTGAAATCTGA
GACAGGTCTCAGTTAATTTAGAAAGTTATTTTGCCACGGTTGAGGACAC
CCACCCATGACAGAGCATCAGGAGTCTTGACCATGTGCTCAGGGTGG
TCTGAGCATCAGCTTGGTTTACACATTTTAGGGAGACATGAGACATCAGT
GAATATATGTAAGATGTACACTGGTTCCTCCAGAAAGGCAGAACAACTT
GAAGCAGGGAGGGAGCTTCCAGGTACAGGTAGGTGAGAGACAAACAATT
GCATTCTTCTGAGTGTCTGATTAGCCTTTCAAAGGAGGCAATCAGATAT
GCATTTATCAGAGTGAGCAGAGGGGTGACTTTGAATAGAATGGGAGGCAG
GTTTGCCCTAAGCAGTTCCCAGCTTGACTTTTCCCTTTAGCTTAGTGATT
TGGAGGCCCAAGATTTATTTTCTTCTACATCACTGTGGGCAGCTGACT
AGGAAAGCTTTGTAGACTGGTGAGGCTGTGAGAGCCAGTGGGGGGTG
GTGGTCTCTGTGCAATGGTAGCAACCACTGTGAGGCTGAGTAACTCAT
TTCCCAACCTCTCTAGCAGCCCCAGTGGAGATACAGAGGAAGCAGACTA
GCGATACAACCCAGCCTGAAGTTTTGTCTGGTGAAGTGAATGGAATAAAA
ATGGGAAGGGTGCTGAAGAGACCAGCAAGAAAATGGTTGAAGAGATGGGG
CACAGAAATTAAGCTGGATCAAAAAGGACGGAAGCAGAAAGGGCCGAT
AGAGAGAGGGGATATCTATGGGTTCCGCGATTCTGAAAAGGACAAATCACT
GGTGCTTTGAGAAGAGAGAGGGTGAGAAAGCAGGAAGGCTGGAGGCTGTC
ATCCAAGAGGCGGACATCTGTGAACATGATTCCAAGAGTCAACAGACCAT
GGGGGTGGCCAAAGGGAGTGCCTCTTCTCACTCTTACTCTTAATTCCTT
GTACTCAAGATAATAAGTTCCCAGAGAGAAGTACCCATATTTAATTCAT
CTGTGTCTTCTAGCAGTACTAAAAATATTATATGAAAGGTATCAAACCT
TTGAGAATGTGTGCTGCTAAATTTGTTAAGGATGCTGGAAAACCTCAAGACG
TCCCTGATCCTGAGCCTGAGTATGAGCCTGTGGTGAGCCCAATGCAGGTC
TCCATTGAGACAAAGGCCTCAGGGAACGGATGAGACCTAGGGACAGAGAT
GCATGCTGGAGCAGCATTTCCCATCTTCTGAGCTCAGGCCAGCTGAC
TGCTTTATGAGTAAACGTTACCAGGGAACACTTTGCACTCTTAACACACA
TGCCACCTGTGACCACTGATCCCTGTTGGGTGACCACTGACATCAGAGA
TTCCGATGGCAGCAATGAAGACAAGGCTATCCTCATTAGGAAGGAAAGGAA
GGAGGAGGGAGGAGGGCAACGAATCTTCTGCTTGTCAACCACGTCCA
TCTCTGTTAGGTGATTTCCCATGTGTGACTTTGTTTATCTTTATAATAAC
TCTGAGAGGTAGGTCTTGATGTCCACATTTTGAACATGAGGACATCCAGC
CAGGAAGTTGAGTTCTGGGACATAGCTGAGAGGGCAAAGCTACATATAA
ACCCCTCTTGTCTTCTGGCTTATCCACTGAGTGCCCCCTGCAATCCA
CCAGCCCATTTGTGAAGTGCACTATAGGTAAGTTGGCACAGGAGGAGT

FIG. 3 (20 of 52)

GGATGTGGGCGATTTTG. JACAGCTCTCCAGGAACCTTACACACTGGTGAG
GAGGGCCAGGTATGTTCCCTGACCAGTCACAATCAAAGCAACCTCCTACTA
ATCAGGGAGGCTTGGTACCTGGGGAATGCTATGTTGAAAGGTTCTTTTCT
GGGTTTTTAAATGATGGGTCTATTTCTTATTCTTAAGATTGCTTTTTTT
CTGGCTAGAACTTAAAAGAAATTTTTCAGTAAATTTCCCTTCCCTGGGCAC
AAAGTGAGCTTGAAATGAATTTCCAGGTGGCCTTGATACTTTAAATATT
GCCTCCTATAAAATCAACCTTTAGAAGAAGGAAGTCAAAGAACATGCTAG
ATTTACAAAAGGTTAATTCCTTGAAATCCAGTTATCTACAGGACAATGTT
GTCAAAGAAAAATTATTTGGCCAGGCACGGCGGCTCATGCCTATAATCC
CAGCACTTTGGGAGGCTGAGGCAGGTGATCACCTGAGGTGAGGAGTTTGA
GACCAGCCTGGCCAACATGGTGAAACCCCATCTCTACTAAAAATACAAAA
AAAATTAGCCAGGTGTGGTGGTGGGCACCTGTAATCCAGCTACACGGGA
GGCTGAGGCAGGAGAATCGCTTGAACCCGGGAGGAGGAAGTTGCAGTGAG
CCAAGTTCAAGCCACTGCACCCAGCCTGGGCAACAGAGCAAGACTTTGT
CTCCAAAAAATTAATCAATGATATTTTAAATTCATGGTAAGGAA
GATTTTCAATCAGAACCAGCACAGAAGATATAGGAAACACTGCAATGGGAC
TTTGCGGTGGGGGAGAGAGATTGAACACAACACTACATATACAGCACGGGCA
AGGACATATTCATAGCCAGGAAGCAGAGCAAAGATCAGTGGATGCGAAAT
TACTAAGAGGAAACATGAAAAATAAGGGAGCTTCTGCCTAAACCCACCTA
ACCGGATCCTTGCTGAAGACAGGACAGGTGATTGGACACCACTTTGGGG
ATGGTGGAGGATGGGGAATCCAGTGAGATTCAAGGGTGATGCGAATTG
AACATACAAAGTTCTTGCTAAAAAAGGATTTTACAAGAAAGTGACAAAT
GTGCTTGGGACAAGGTGCAGGAGCCCGACGGAGATGTGGTCCAGCAGAGA
ATATGTGCGAGATGATAGGTGAGTTCTCTGACGAAGGATATATGCTGAT
CCAGCCAGGGTGAAATGCTCAGAGAAAGCACGGAGGGGCTATGTCCGTTG
CCCCAGTCTCCACGGGTCAAATCTGATCCCGTTGTGAGTGTGGCCGTTT
GTAGAAAGCAATCAGGGGGGTCCCTCCCC

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AATATATATTTTTTATANNATNTGAGACAGGTTCTCACTAGGTTGCCAG
GCTGGTCTTGAATTCCTGCCTTCAAGTGACTCTCCACCTTAGCCTACTG
CATAGCTGGGATTACAGGCACAAACCACTGCATGCAGCTAACTTTGCTTC
TCATTCAGCACTTTTTATTCCACTGATTATATGTATATGTATATCTGCA
TCATCTCTCTCTCTCTCTCTCTCTCTCTCTATATATATATATATAT
ATGGAAATATCTCTCTCTCTCTCTATATATATATATGGAATATATATCT
CAGTCTCTCCTATCCTTTAATCAGTTTTGCTATCCTGTCAATCCCC
CAACGAGTGTGATGTTGTGAAATATATATTTGTTCTTCATCTCCTGTTTC
CTGACATACAGCTTTTAAAAACCCCTGGAATCTCTGGAATAATAAGAGTG
TCTTTTGCATGCTAATAGATGACTGCTGGCTGGCAGCCCCAATGCAGTAG
CTTCATGATGGGGTTTGTACAGGAAAGACCAAGGCAGGATTGGAGACTT
GAGACTGTTAGCCCCACTCCCCAACCACTGGAGGGAGTGGAGGGGCTGAA
GGTTGTGTGAGTCACCAATGGCCAATGGTTCGGTCAATCATGTGTATGTA
ATAAAGCCACTCTTAAAAACCCAAAAAGGACAGGGTTTGAAGGGCTCCC
AGATAGCTGGACACATGAAGGTTTCTGGAGGGTGGTGGCCAGAGGGGCA
TGGAAGCTCCACACCCCTTCTCACATGCTTTGCTCTGCGCATCTCTCAT
CTGGTGTTCATCTGTATCCTTTGTAATATCTTTAGAATAAACTGGTAAA
CTTAAGTGTTTTCTGAGTTCTGTGAGCTGCTCTAGCAAATTCACGGAAC
CCGAGGGAAGCAAACCCAGATTTATAGCCATCAGTCAGAAGCATAGGTGA
CAACCTACCACCTTGTAACTGGCACCTGAAGTGGGAGGCAGTCTTGTGAGA
CTGAGCCCTCAACCTGTGGGATCTAACGCTAACTCCAGGTAGATAGTGT
GGAGTGAATTAGGACACCCAACTGGTGTGCGCTGCTGGAGGACTAGTGGT
GGGAGAAAATCCCCAAGCATTTCCGGTGACTAGAGGTACAGAAGAACTCAG
TGTTGAGGTGTTGTGACAGTATGGTAGGGAAAACCTGCGTCTGGTTTTTTC
CTTTTACAAATCAGTTAAATATTTAACACAAGTCTACTGTATATTAGTAAA
AGGGTTACATTTTTTAAATGTCTTGACAGTTGCACTTTGACAACCTCCATA
TCAATCACTTTTTTTCGTGTCCGTTTGAACCAAATCACTTGGGATACC
ATGAACAGGCTGCAGCGTATTTCCCAAGGCCTTGAAAGCTTGGAGGCCAT
TTTGGCAGCCNTAAATCCCTGTGAATACCAGGCTTCGTGGATTTAAAAAT
AGACTTGAGGCCAGGCCCTGGTGGCTCACACCTGTAAGCCCAGCACTTGG
GAGGCAGAGGCCGATAGATCACAAGGTTAGGAGTTGAGAGCCAGCGTGGC
CAACATGGTGAAACCCGCTCTCTACTAAATATACAAAAAAATTAGCCG

FIG. 3 (21 of 52)

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GGCGTGATGTTACACG CAGTAGTGCCAGATACTCAGGAGGCTGAGG CAG
CAGAAATACTTGAACCTGGGAGGCAGAGGTTGAAATGAGTCAAGATCGTG
CCACTGCACTCCAGCTTGGGCGACAGAGTGAGACTCAGTTTTTCAGGGGAG
TTAAACAATAACAAAAAAGAAAAAGACTTGAACAATGAGGCTCCACTGG
ATGGATTTAGGGGAATTACAGGAAGCAGGACCTGACGGTGCAATGCCACA
CTCCACCTGTCCAGAATTGGACCTCACCAGGGAGGTCTGTGGGGACAGG
GAGAGGCCCTCTGCCTCCACCCCCCTCTCTACTCCCCAACCTTGAGTCA
GGCTGAATGTAGTAAACCTGGAACAGAAAAGTT CAGTTTGGCAATAGGTA
TCTGAAGGACTCCAGGTGCTTCTCCCTTGATTCAAAATTTTACTTATAAA
AAAAATTATAAGAAAATTCTACTTAAAAGAAATAATCAGGGAGGTACAAC
AAATTGTACTTTTTTTTTTTTTTTTTTTTTTGTAAATGGAGTCTCACTG
TTGCCCATGCTGGAGTACAGTAGTGTGATCTCGGCTCACTGCAACCTCCG
CCTCCTAGGTTCAAGTGATTTCCTACTTCAGCCTCCCAAGTAGCTGCGA
TTACAGGTGTGTGCCACCACACCCCGCTAATTTTGTATTTTGGTAGAG
ACGGGGTTTACCATGTTAACCAAGATGGTCTCGAACTCCTGACCTCAGG
TGACCCACCTGCCTCAGACTCCCAAAGTGTTGGGATTACAGGGGTGAGCC
ACTAAGCCCAGCCATTGTACATATTTTGTGGGTATTTACTAAAACATTAT
TCAAAATAGTAAAAAAAATTGAAATAAACTGGGGACTGGTTAAATAATT
TTGGGTACAACCACATGATGGAATACTATACAGCCATTAAAAATTACATT
GAGGCCAGGTGTGGTGGCTCATGCTTGTAACTTAGCACTTTGGGAGGCC
AAAGTGGGAGGATTGCTTGGACCCAGGAGCTCAAGACCAGCTTGGGCAAT
GTGGCAAAACCTGTCTCTAAAAAAAATAACAAAAAATTAAAAAGCT
GGGTGTGGAGGCACACACCTCTAGTCCAGCTACTCAAAGGGCTAAGGTG
GGAAGATCACTTGAACCGGGGAGGTCAAGGCTGCAGTGACCCAAAATCGG
GTCAATTGCACTCCAGCCTGGGCAACAAAGCAAGACCCTGTCTCAAAAAA
AAAAAATACATTGAAGAATATCTTACGGTATGGATAAATATTCAATTTA
CAGTGATAGATGCAAATAAAGCAAATTACAAAATATACAGTTTAATTCC
AACTTTGATACTACATATGTATATATGAATACATGCATATGTTATGTATG
TATATGTAATATAACAATATATGTTCTATATATGGATATTATATATTTA
CACATACATACACATATATAATATCTTCTCTAGAGAGCAGAAAGAGAG
TAGACAGATAATGAAGATAGGATACAACTCCAGTCCAGCTCAACCTAGGG
GACTTGTTTTTAAAGCCTCAGGAGAGAGAAGTTGGGACTAGAAAGCAAGGC
AGCTATTTGTAAGCATCTTTGTGTTTCATGCTATTGGGGTGGGAAACAAC
AGCACAACTTTTGAAGGCCCTTTCTACTCACCCCAAACTGCAGAGCA
GCTTTAGGACCCTCAGAGTTCAAGAAGACCATTTCAGAGTAGAAGAAGT
AAAAACATGTATGAACCTGACCCCTGAGCTCATGGACTGTGCCATGAGGGA
AATTCTTAAACACAGCAGGAGAGGCCCTGGAGGAAGGCAGAGGCCCTGCAT
CAGCAAGTCCAGGCAAAAGCCTGCATTCCATAGATGCTCATCTCTCTGGC
TGGTGAGGTCTAAAGACGTTTGGTCTCAATATTAAGTCTCTGAGAGAGG
TCACAAACCCAGTCCCTTGGCCACAAAAGGAAATAAATCTGGCTTGAGA
CATTAGGGAGGAACAGGGCAAGGGGAGGTTCAAGAAAGTTTTAATGGATG
AGATGATATTTAAGCAAGGCCCTGGAAAATGAGAAATTCACCAATAGCC
ATATGGTAGGTGAGAAAGCAAGATAAGGAGGGGGCAAGTGCAAGGGGCA
ACATCAGATATGACCAGGGTGTCTGTTGGGCAATGGCTGATGGAGAAGA
TTAGACTGGAGTTTGGGAATGCCACAGTATCGAGGTTGGATTTAATCCTA
TGGGTAATAAAGCCAACTGTTCAACCCCCAACCCACTTGCAATATGGCTC
CAAAATAGCAGGTGTTTGTATAAATGACTACTTTTACTCTACTATTCCCT
CCCTCTTAAGAAGAAAAAGAAAGTGGAGGCTCAGAGAAAGGCAGTGGCTT
GTCCCAATCACACTATGATTTGGCCACAAAACAAGAACGAAATGTTACAC
CCAAAAATGCTGCCTCCACCTCCCTTCTTGCTTTCTCCCTGCTGGACT
ACAGACTATCTCAAGAGTGACGTACACCATCAGGGCTTCAGCTTTTCCCC
GAAACAAATGCCAAATATTAGCCATACGTCACTGTAGTAAGAGCCCTGAA
TTGGGAATCCAGCTTTGACGCAGACATGCTGATTGACTCTGTGACCATT
CTCTTCACTTCTCCACTCTATTCTTCCCCACCTGTAAAGTGAGGTCCTT
CCAGTTATAAAAAACAGATGATGCTATTGTCTGTTTTGTATCTAATCTTG
CTGTGTTATAAAAAAAAATAAGGCTCTGTACATTCACTTGGCCAATTC
CCTTCTTATCTCTACTTCCCACAGCCCCCTTTTCTACAGAAAACCAGCAT
TGTCTTCTGGATCCATCTCTTAAGAAAGCGCTTTGCTCCCGGTTATT
TAGGTGATAAGAAGTGTCTTAGATGACAGCCCTGGAATGGGCTGGAGGCA
ACAAAAAGCAAGTGAAATAGACAGTTACAGCGACGACAATAATAACAAC

FIG. 3 (22 of 52)

CAACACCTCTCACTAAAGAGAAAGAAATAAAAAAGAAAATTAAAAATCTGC
CGCAATGCCCAACAGTCATTGAATAACTGCATGTGTACAGCACTTGGTT
ACTTTTACATACTTCATATTTTAGCCTTCATAGCAGCTCACAGGGGTGGA
TTTAATTTTAGTCCAACTCCTGTACGGTGCCTGGCACAAGTATAATAA
ATGTTCTGTGAATAAATGACCCTCTTTTAGATGAGGAAATCGAGGCTCA
AGGAGAACAAGCAATGTAATGTCCCCCTCCTGTTGAGCCATCTGCCTTTC
AGGCCACTGAATGCAGTAGTCCTCAGTGCCCTGAACTTGACCCTCTTCTG
CTTTTCGGACTGGTCTTCTAATCCCGTTGTGACTCACTACACCACCTCT
CCTGCATATGACATCTACATTTTAAAAACAAACCGTATGGAAATAACACAT
TAGTGGGCTTGTTCCTCCACCCCGCAAAAAAAGGCCTCTTTATAACA
GAACTTCTCAGGCTGGTAGGGGAATTTTATCCCCCATTTATGGTAGAA
AGGCCCTAACCTTGGACCTCAGCCATAGCTATTACATGGGGGAATGAT
GAATAACATGGGGAGCAGCATGTAAATATCATTGAGCCGTAGTCCAGACC
TATAACATC

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GGGGGAGCTGCATGTGCCTGTGCGAGATCTGGGGGAGGAACAGGAAGATCA
AGAGTTCTGTGTAGGACATGTTAAGTTGAAGGTGCTTACAGGATAGCCAG
ATGAAGCATCAGGTGTGCAGTCAAAGATATGAGTCTGGAGCAGCACATCC
TAAGTCACCTCCTGCACCAACACAGAACTCCAGGCCACTCACTTGAGCT
CTCCCAAATAGTTTCCAAGTGTCAATTATGTTAATAACCTATGAGCTTGAA
CACCAGATTCAAACCCCACTGCATGGCTTTTAAAGACCATCTCAAGGGCT
TGACACTCCAGGGAGCCAACTAAAGATGCCTGGTCTTACCATCAACCTCC
ACCCCATTTTTTATAGAAAATGTTTCTACCTGTCTAAGGCAGGGTCTTG
CCCCACTCCAGGCCCTTTAGATCCCCAATATTCTCTCTCCCTGAACCA
AAACCTTCATCATCTTCCAGCATGGGTGGGGCTCCATTCTTGCTTCTGC
TCCCTGAGCAGAAGCAAGTTTCTCCCACTTGACCTGATTCTCTCTCTA
AGTACCAGTCACTGCTTGTGTTCTGGAATGAGAGAAAAAGACAGAGTGAG
AGAGACAATCCAGAACTCTTGCTCACTCAGCTAGGCTGGGCATCTGGG
AGGATGGCTGTGTCCATGGGAACCTGGGAAAAGCCACACCTTGGCACCC
TGGTCACCCACCTGTCTCCCTGGCAGATTCCGCACTGCTCTCTTGACCC
TCTACCAGGGCTAACCGGCCTGCTCACTCTCCCAAGCATGTCTTCCACG
CCCCTCTCTAATTATTACATTTCCCTTCACATAAACTGCCCTTCTCTCC
AATCACCACATGTTCACTTCCCAACCCAGCTGTCAAAGTCTGGCTCAACCT
CATTCTTGAAAAGGAAAAAACAACAAACAACAACAACAACAAGCAAAAA
ACCTATGATGGATTAAGAACACACTTCATTCCAGGAACATGCTTATCTCC
CTTAACCTTCACAACAACACTACAGCAGGTAGGTGTTATCACACCCATCTCT
CAGGTGAGAAAACAGGCTCAACGAGTGCAGGAGGACACAGCAAGTCAGTG
ACAAAGCTTAAATTCAAGCCCCAAGCCTGTTGGCAACCAACGTCTGTACCC
TTGATAGCTACCTCATTTACACCAAATCCAGTGGCCTCAGGCCCTGGCTG
CACACTGGGATCACCTGGTGCCAGACCACATCTTAGACCAGTCATACAG
AATCTCTTGGGCTGGGATCCTCCACGGTACATTTTAAGGGTCCCAGGTG
AGTTCCACCATGGACCCAGAATTGAGGACCCAATACCGTATACCATCTCC
TTCTTCATCTCTTCTAAGGCATCTCTTACTCGCTGTGCACTCCCATACCA
CTTTGTTCAATCATCCAATCATTCAATTGAGTCAGTTAGTCAGGAGC
TACTCACTAGTCCCCTGCCAGGTCTAGTCATGACATAGGGCTCTGGGGA
CCAACAAGAAGCAGGACCCATGCCTCCTGCTCTCATGGAGCTTGCTCTGC
AGCAGAGGAAGCAGTCAGTGAGATGTAGCAAATGTGAAATGTGCACAGAT
GGGAAAAGCAAACTTTAAACTTTTAGGACAAAATACACAAGAAATCTT
TGCAACTTTGGGACAGGAAGGAACAACATTCCTTACACATGACACCAAAG
GAATCAACCATAAATAAAAAGGTGATCAATTTGACCTCATTTAAGTGTTA
AGCTTTTTTCATTGAGAGACACCAATTAATAAATAAATAATACATGCCACAA
ACTGGGATACAATAATTTACAACACTTATGTCTCACAAGGATTAGTTTTTC
AGAATATATAAAGAACTCCCGGCCGGGTATGGCCGCGCACGCTGGAATCT
CAGCACTTTGGGAGGCCAGCGGATCACATGAGGTCAGGAGTTCAAGACCA
GCCTGGCCAACATGGCAAACTCCGTCTCTACTAAAAATACAAAAATTAG
CCAGGCATGGTGGCGGGCGCCTGTAATCCAGCTACTCAGGAACTGAGG
CAGGAGAATCACTTGAGCCCAGAAAACAGAAGTTGCAGTGAGCTGAGCTC
ACATCACTGTAAGCCTCGGTGACAGAGTAAGACTGTCAAAAAACGAAAA
CAAAAAACAAAACTCCTACAAATAAATAAGAAAAAATAGCCCAGCAGGA
AAAAGTATATACATTTCTAAAAAGAATAAATACATTCTGTCACTTTTCTA

ACATATATTTTTTAAGAGTAAATACAAATGGTTAGGAAACATTTTTTAAA
ATGCCCAACCTCATTAAAAATTATAGAAGTGAATAAAGCCACAATAAG
ATACGATTTTATACCAAATACAGTGTCAACACTTTGCAAGTCTGACCTCA
CCAAGTCTTACCAGACGTGTGCACTGACGTGGCTGCTGAGATACTGATGG
TGGGTCTGTAAATCTGTACTACAAACAATTGCAATAAAATGTAATAAATA
TACAATAGGTGGAGCAGGAAGTGACCTGCAACCATATAGCAGATAGGGCA
GGAAAAAGCCTATGAAAGCTGACATCAAAGGGATAAGTTCAGTTACCCA
GCTGAAGGGAAGGAGGGTGTTCAGATAGAGGAAGGATAAGCATGACCTA
TTCAAGGCCAGTGAAAGAAGCGTGCAACGGCCAGTCAGGAGAACCTGAA
ATTGTGTCAAAGAGCTTGGATGCAAAGAGCCGTGGGAGACTATTGGGGGT
TTTAAGCAGGGATATAATATTCAATCAAGCATGCAGTAAAAGGTCACTGG
CACCTGCCATGGGCCAGGACTCGGGCTCTACATGATTGCGTCTGTTTTGG
AAATATCACCTCGGCTGTGAGATGAAGAACAGGTAGGAGGGTCACAAAC
TTGAAGCAGAGAGACTGTTGAGGAAGTAAGCTGTTTTTGTGTGGACTGTG
GCAATCACAGAGGCAGAGGATATAAATGCACAGAGACACAAGGCATGTGG
GAGGCAGAAGGAATCAAATACAATGAGTGATCAGATGTGGGGTTAGAGTG
GTGAGTGAGAAACATACTCAAGGTGACACGCCAGGTATCTGGGTGGAT
GGTAAGACATTCATGGACTAGGATCGAGGAANGAGGTGGGGAATGGGACC
ATACCTGCAGTTTATAAGGGGTGGACGAGGGAAGATTATGCGGGAGACTG
AGAGAGGAATAGACAAAGGAATCCCGGTGCAGTATTACAGAACTGGGGT
GGGAGGGGGTGTANTTCAAAGGAAAGAAAATTGTCAAATAGTATGAA
ATGCTGCAGAGAACTCACGGATTTTTTTTTTAAGCTTAGAATTATTATCAT
TGACTATGTGAATAAGAATAACTTTTATGAAAGAAGTTTTGCTTAAGTAG
TAGGAAGAAGCAAAATTGTTGAGGGCTGATGAGTGGGAGGAGAAGTAATT
GAAGGCACTCTTCAAGAGAAACAAAGCAGAAGGTGAGGAGAATACTAAT
GAAGGAGTTACGGCCTTCACTATTTTTGTTTTGCTTAGATAAGCAAGACT
TGAGTGGGTCTGGTGAGGAGAAACAAGTAGAGTACAAAGTTAAAGGAGAG
ACAGACAGAGATAGAGATAGGGACAGAGAGAGAGACAGAGACAGAGCACA
AAAGAGCAAGGTCCCTGAGAACACGGGCCTTCTGTTTAAACCCAGCCAG
ATGTATTGCAATTCAATTCCAGTACTAACCACCCAGAGTTTGTGTAGACT
CTACAAGTTAAAGAGCATGGTCCCAACAGACTGCTTCTACGTCAGATG
CCAGGCACACTTCAGGGGTCCCCAAGCCACTCATGTTTTTTGAATGACTG
CCATAAGTTCAAAAATTCCCACAATTCTCTCAGATTCAATAACTGGGTAT
AACCACCTCATAGAACTCAAGAAAATGCTATCATTATTATTACAATTTTAT
TATAAAGGATACAAATCAGAAGGACTAGCCAAATGAGGAGACACATAGAG
AGAGGACTAGTAAAAAACAGAGCTTCTGCGTCCTACCTTCAAGGAATCAG
GATGCACCAACCTCCAGCACATCAAGTGCTCATCAACCAGGAAGTTCCT
CTGAGCTCCAATGTCCAGAGATTTTAGGGAGGATTATTACATAGGTATC
ATTGATTAAATCATTTGGCCATGTACTTGAACCTCAATCTCCAGTGTCCCTC
TTCTCCCTAGAGGTCTGAAGGGTTGGCTAATATCATGTGGCTCAAAGCCC
CAACTCTAATTACCTTTTTTGGTCTTTTCAGGGACTAGACCCCATCCTGAA
GCTATCTACAGGCCCTGCCATGAGTTAGCTCATTAAACATAACAAAGACAC
TTATATTACTCAGAAAATTCCAACAGTTTTAGAAGCTCCATGTGAGGAAC
CTGGGACATAGATCAAATCTTTTTTTTTTTTTTTTTTGGAGACAGGGT
CTTGCTGTGTTGCCAGGCTAGAGTGCAAGGACAGATCACAGCTCAATGC
AGCTTCAACTTCCAGGCTTAAGTGACCTTTCCACCTTAACCTTCCAAGT
ATCTGGGACCACAGAAAATGGCTAATTATCCTGGCTGATTTTTAACTTT
TTTTTTTTGTAGGGATGGGATCGCCCTGTGTTGCCAAGGTTGGTCTCAAA
CTCCTGGGTTCAAGCAATCATTCTGCCCTGGCTCTGTGATGGTTAATAC
TGAGTGTCAACTTGATTGGATTGAAGGATACAAAGTATTATTTTGGGTG
TGTCTGTGAGGGTGTGCCAAGGAGATTACATTTGAGTCAGTGGACTGG
GAAAGTCCACCTTTCCAGTGACTGGGAGACCCACCTTCAATCCAGGT
AAACACAATCTAATCAGCTGCCAGTGTGGTCAGAAATAAAGGAGGCAGAA
GAACAGGGAACACTAGACTGGCTTAGTCTTCCAGCCTACATCTTTCTCT
CATGCTGAATGCTTCTACCTCGAACATCAGCCTCCAAGTTCTTCAGTT
TTTGGACTCTTGACCTTCAACCACAGATTGAAGACTGCAGTGTGGCTT
CCCTGTTTTTGAAGTTTGGGACTCAGACTGGCTTCTTGCTCCTCAGCT
TGCAGATGGCCAAATTGTGGGACTTTAACTTGTGATCATGTGAGTCAATAT
TCCTTAATAAACTCAGATATATATATATGTATCAGACATATATATATATC
CTATTGTATATTATATACAGATATATAATATCCTATTATATACAGATATA

FIG. 3 (24 of 52)

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TAATATCCTATTATATATACAGGTATATATATATATATGTATCATATATA
TATCCTATTGGTTCTATCCCTCTTGAGAACTCTGACTAATACAGCCCTCCC
AAAATGCTGAGATTACAGGAGTGAGCCACAGCCACCATGCCAGCCCCAA
ATTCTTAATTATACAACATGGGTCCAGAGATCAGGGCCTGGGTAGGATG
CAGCAATAAGAAAAACAGATGGTGGATGGGGACACATGTTGGAAGTGTGGC
AGGACATGGCTGAGGGAACTCATAGGATGGTGTCTATTTTCATGGCTGAG
TGTGAGGAACAGCATAAAGGTCAAAATTTTCAGGTCAATGGTGAGTTTTTA
AATTGTTGCTGTGAACCCCAAAATCTGACCCAGGTCTCAGTTAATTAG
AAAGTCTATTTTTCCAAGGTTGAGAACACCCACCCACTCACGACAAGAGC
ATCAGGAGGTCTGACCACATGTGCCCAAGGTGGTAAGAGCACAGCTTGG
TTTTATATATTTTAGGGAGACGTAAGTCATCAATCAATATATGTAAAGATG
TACACTGGTCTGCTAGAAAGGCAGGACAACTGAAGCAGGGAGGGGGC
TTCCATGTACAGGTAGGTGAGAGACAAACAGTTGCATTCTTTGAGTTTC
TGATTATCCTTTCCAAAGGAGGCAATCAGATGTGCAATTATCTCAGTGAG
CAGAGGGATGACTTTTGAATAGAAAGACAGGCAGGTTTGCCCTAAGAAGTT
CCCAGCTTGACTTTTCTTTAGCTTTGTGATTGGAGGCGCCAAGATTT
ATTTTCTTTTCAATTTCCCCCTTTCTTTTAAAGATCTTTAAAGAA
AGCTTTTAAAGAAAGAAATGAGTCTCTGGTCCCAGGTTTCATCTGAATTCT
CGAGGGGAGGATGGTTTATCCTAAACGGGTGGTTCTGAATTTTGAGAAAG
TGCATTGTAC
>Contig32
AAAAGCCATACGAATGAGGAAGAATTAAGGGCCAGAACAAAAACAAGAAGA
TGAGGGAAAGTTTGGAACTTTCTAGAGACTGGCTAAATGGTTGTGACCAA
AATGCTGATAGTGATACGGACAATGAAGTCCAGGGTGACAAAGTCTCAGA
TGGAAATGGGAATTTGTTGGAACTGGGCAAGGTCAACCTTGCTATGA
CTCAGCAAAGAAATTTGGGTGCATTGTGTTTATGTCCTGGGGATCTGTGGA
AGTTTGAATGTAAGAGTGATGACTTACGGTAGGGTATCTAGTGAAGAAA
CCTCTAAGCAACAAAGTGTGTTGCTTAGAAATTTCTTTCTTTCTTTTTT
TTTTTTTTTGAGCTGGAGTTTTGCTGTGTGCGCCAGGCTGGAGCGCAGTG
GCGCAATCTTGGCTCACTTCAAGCTCTGTCTCTGGGTTTATGCCATTCT
CCTGCCTCAGCCTCCCAAGTAGCTGGGACTACAGGCGCTGCCACCATAC
CTGGCTAATTTTTTAGTATTTTAGTAGAGAGGAGTTTACCATGTTAGC
CAAGATGGTCTCAATCTTCTGACCTCGTGATCCACCCGCTTGGCCTCCC
AAAATGCTGGGGTTACAAGCATGAGCCACCCGCTTGGCCTGCTTAGAAA
TTTCTAAGCCAGGATATGGCCTGTCTGCTTCTAACAGCCTGTGCTCAGGG
GTAAAGAAATGACTTAAAGTTGGAACCTATGTTTAAATGGAAGTAGAGT
CTAAAAATTTGGAAAATTTGCAGCCTGGCCTTGTGGCAGAGAAAGAATCC
AAGTAGGCTGCAGAGCAATCATTGCTAGAGAGATTAGCATGACTAAAAGG
GAGCCAAGTGCTAATATTCAAGACAATGTTTAAAGGCTTGGGGCATT
TCAGAGATCTATGAAGCAGCCCTCCCATCACAGGTGCAGAGGTTTGGTG
CACTAGGCCCAGAGGTTTTATGGGCCANNGCCAGGGCCACACTGCTATGC
ACAGCTTTGGGACACTGCTGCCCGCATCCAGGCCACTCTGCTCTGGCTCC
ACCTTGGCTCAAACGGGCCAAGATAGAGCTTGGACCACTGCTCCCGAGG
GCACAAGCCATAAGCCTTGGTGGTTTCCATGTGGTGTAAAGCCTGCAGGT
GCCCAGAATGCAAGATTGAGGGAGCTTGGGCACTTCCACCTAAATTTAG
AGGATGTGTGAGAAACCTAGGTTCCAGGCAGAAGCATGATACAGGGGC
AGAGCCCTTGACAGAGAACCTCTACTAGGGCAATGCCAAAGGAAAATGTGG
GGTTGGAGTCTCACACATGGTCCCCACTGGGGCACTACCTGGTGATACT
GTGGGAATGGGGCTGCTGCCCTCCAGACCCAGAAATGGTAGATGCACTGG
CAGCTGGCACCCCTGAGCCTGGAAAAGCTGCAGGCACTCAACTCCAACCCA
TGAGATCAGCCACATGGGCTACTCCAGGGAAGCCACAGAGGCAGGGCT
GTCTAAGGCCTTGGGAGCCTACCCCTTGAACCAGCTTGCAGGACATGGAA
TCAAAGATTATGTTGCAGCTTTAAGGCTTAATGTTTTCCCTGTCAATTT
AGGCTTGTGTGGGACCTGTTGCTTTTTTTTTTTTTTTTTTTTTTTTGGT
CACAGGTGTTTGAACCAGAACATTCATCTTGAATAGGGGCTGGGTAAA
ATAAGGCTGAGACCTACTGAGCTGCATTCTAGGAGGTTAGGAATTTCTAA
GTCACAGGAGGAGATAGGAGGTCCGCACAAGATACAGGTAGCGAAGACCT
CGCTGATAAAATAAGTTGCAGTAAAGAAGCCAGCCAAAACCTCACAAAGCC
AAAATGGTGATATGGTTTGGCTCTATGTCCCCACCCAAATCTCATCTCAA
ATTATAATCCCATAATCCCCACATGTTGAGGGGAGGACCTGGTTGGAGG

FIG. 3 (25 of 52)

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TGATTGGATTATGGAGGCAATTTCCCCCATGCTGTTCTGGTGATACTGAG
TGAGTTCTCATAAGATCTAATGGTTTTATAAGTGTGGGAAGTTCTCCT
ACACACATGCTCACACTCTCTCCTGCAGCTTTATGAAGAAGGTACTTGCT
TTCCCTTCTGCCATGATTGTAAGTTTCTGAGGCTTCCAGCTATGCAGA
ACTGTGAGTCAATTAACCCGTTTTCTTTATACATTACCAGTCTTGGGCA
TTCTTTACAGCAGTGTGAGAACTGCTGGCGATGAGAGTGACCTCTGGTT
GTCTCAGTCTCATTATATGCTAATTATAATGTATTAGCATGCCAAAAG
ACACTCCCAACATGACCCCAACAGTCATGCCTGTGCCGGTCTCAGCACCA
TGACAGTTTACAGATGGCATAGCAACGTCTAAAAGGTACCCCATATGGAC
TAACAAGGGGAGGAACCCCTCAGCTCTGGGAAGTGCCTACCTCGTTCCAG
AAAGCTTGTGAATAATCCACTGCTTGTTTAAACATATAATTAAGAAATAAC
TATTAAGCATCCTTAGTTTACAGCAGCCCAAGCTGCTGTTCTGCCTATGGAG
TAGCCATTCTTTATTCGGTTACTTTCTTAATAAAATTGCTTTTACTTTAC
TGTATGTAATCGCTGGAATTCTTTCTTGACGAGGTCCAGAGCCCTCTC
TTGGGTCTGATCGGGACCCCTTTCTGGTAACATTTTGACCAATTTCTCC
CTTCTGGAATGGGAATGTTTACACAATGACTGTATCACTTTGAATCTTG
GAAGTAAATAATTTGTTTTTACTTTTACAGCCTCATAGGTGGAAGGAAC
TGACTTGAATTTTACAGATGAGACTTTGGACTTTGGGACTTTTGGGTTGGGG
CTGGAATGAGTTAAAAGTTGGGGGGATTATTGGGAAGGCACGATTTTATT
TTGCAATATGAGAAGCACATGAGATTGGGGGACCAAGGGTGAATAATA
TGGTTTGGATGTTTGGCCCCCTCCAAATCTCACATTGAAATGTAATCCCA
GTGTTGAAGTGAGGCTGCTGGAAAATGTTTGGATTACAAGGCTGTGAG
CACATTGGATAAGACGTGTAGGNCCC

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CCGAGCTCGCTGGTTAATTCTGTGGCTCCTGTGACCCTATTATAGCACC
AGGTCTATGACCAGGAGAATTAGACTGGCATTAAATCAGAATAAGAGATT
TTGCACCTGCAATAGACCTTATGACACCTAACCAACCCCATTTTACAA
TTAAACAGCAACAGAGGGAATACTTTATCCAACCTCACACAAGCTGCTTTC
CTCCCAGATGCTGCTTTTGGCGTTTATTATTTTATAGAGATGGGGGCT
TCACTATGTTGCCACACTGGACTAAAACCTCTGGGCCTCAAGTGATTGTC
CTGCCCTCAGCCTCCTGAATAGCTGGGACTACAGGGGCATGCCATCACACC
TAGTTCATTCTCTATTTAAAATATACATGGCTTAAACTCCAACCTGGGA
ACCCAAAACATTCATTTGCTAAGAGTCTGGTGTCTTACCACCTGAACCTAG
GCTGGCCACAGGAATTATAAAAGCTGAGAAATCTTTAATAATAGTAACC
AGGCAACACCATTTGAAGGCTCATATGTAAAATCCATGCCTTCCCTTCTC
CCAATCTCCATTCCTTCCAACTTAGCCACTGGCTTCTGGCTGAGGCCTTAGC
CATACCTCCCGGGGCTTGACACACCTTCTTCTACAGAAGACACACCTTG
GGCATATCTTACAGAAGACCAGGCTTCTCTCTGGTCTTGGTAGAGGGCT
ACTTTACTTTAAACAGGGCCAGGGTGGAGAATTCTCTCCTGAAGCTCCATC
CCCTCTATAGGAAATGTGTTGACAATATTGAGAAGAGTAGGAGGATCAAG
ACTTCTTTGTGCTCAAATACCACTGTTCTCTTCTTACCCTGCCCTAACC
AGGAGCTTGTCAACCCAACTCTGAGGTGATTTATGCCTTAATCAAGCAA
ACTTCCCTCTTCAGAAAAGATGGCTCATTTTCCCTCAAAAGTTGCCAGGA
GCTGCCAAGTATTCTGCCAATTCACCTGGAGCAATCAACAAATTCAG
CCAGAACACAACCTACAGCTACTATTAGAATATTATTATTAATAAATTC
TCTCCAAATCTAGCCCCCTTGACTTCGGATTCACGATTTCTCCCTTCTC
CTAGAAACTTGATAAGTTTCCCGCGCTTCCCTTTTCTAAGACTACATGT
TTGTCTATCTATAAAGCAAAGGGGTGAATAAATGAACCAAAATCAATAACT
TCTGGAATATCTGCAACAACAATAATATCAGCTATGCCATCTTCACTA
TTTTAGCCAGTATCGAGTTGAATGAACATAGAAAAATACAAAACCTGAATT
CTTCCCTGTAAATTTCCCGTTTTGACGACGCACTGTAGCCACGTAGCCA
CGCCTACTTTAAGACAATTACAAAAGGCGAAGAAGACTGACTCAGGCTTAA
GCTGCCAGCCAGAGAGGGAGTCATTTTATTGGCGTTTGAAGTCAAGCAAGG
TATTGTCTTACATCTCTGGCTATTAAAGTATTTTCTGTTGTTGTTTTTC
TCTTTGGCTGTTTCTCTCACATTGCCTTCTCTAAAGCTACAGCCTCTCC
TTTCTTTCTTGTCCCTCCCTGGTTTGGTATGTGACCTAGAATTACAGTC
AGATTTCAAGAAATGATTCTCTCATTTTGTCTGATAAGGACTGATTGTTTT
TACTGAGGGACGGCAGAACTAGTTTCTATGAGGGCATGGGTGAATACAA
CTGAGGCTTCTCATGGGAGGGAATCTCTACTATCCAAATTTATTAGGAGA
AAATTGAAAATTTCCAACCTCTGTCTCTCTTACCTCTGTGTAAGGCAAA

FIG. 3 (26 of 52)

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TACCTTATCTCTGTGGTGTTTGTGTAACCTCTTCAAACCTTTCATTGATTG
AATGCTGTCTCTGGCAATACATTAGGTTGGGCACATAAGGAATACCAACA
TAAATAAAACATTCTAAAAGAAGTTTACGATCTAATAAAGGAGACAGGTA
CATAGCAAACCTAATTCAAAGGAGCTAGAAGATGGAGAAAATGCTGAATGT
GGACTAAGTCATTCAACAAAGTTTTCAGGAAGCACAAGAGGAGGGGCTC
CCTCACAGATATCTGGATTAGAGGCTGGCTGAGCTGATGGTGGCTGGTG
TCTCTGTGTGCAAAAGTCAAGATGGCCAAAGTTCCAGACATGTTTGAAGA
CCTGAAGAACTGTTACAGGTAAGGAATAAGATTTATCTCTTGTGATTTAA
TGAGGGTTTCAAGGCTCACCAAAATCCAGCTAGGCATAACAGTGGCCAGC
ATGGGGGCAGGCCGGCAGAGGTTGTAAAGATGTGTACTAGTCCTGAAGTC
AGAGCAGGTTTCAGAGAAGACCCAGAAAACTAAGCATTTCAGCATGTTAAA
CTGAGATTACATTGGCAGGGAGACCGCCATTTAGAAAAATTATTTTGA
GGTCTGCTGAGCCCTACATGAATATCAGCATCACTTAGACACAGCCTCT
GTTGAGATCACATGCCCTGATATAAGAATGGGTTTTACTGGTCCATTCTC
AGGAAAACTTGATCTCATTTCAGGAACAGGAAATGGCTCCACAGCAAGCTG
GGCATGTGAACTCACATATGCAGGCAAATCTCACTCAGATGTFAGAAGAAA
GGTAAATGAACACAAAGATAAAATTACGGAACATATTAACTAACATGAT
GTTTCCATTATCTGTAGTAAATACTAACACAACTAGGCTGTCAAAATTT
TGCTTGGATATTTTACTAAGTATAAATTATGAAATCTGTTTTAGTGAATA
CATGAAAGTAATGTGTAACATATAATCTATTTGGTTAAAAATAAAAGGAA
GTGCTTCAAAACCTTCTTTTCTTAAAGGAGCTTAACATTCTTCCCTGA
ACTTCAATTAAAGCTCTTCAATTTGTTAGCCAACTTCAATTTTACAGAT
AAAGCACAGGTAAAGCTCAAAGCCTGTCTTGATGACTACTAATTCAGAT
TAGTAAGATATGAATTACTCTACCTATGTGTATGTGTAGAAGTCCCTAAA
TTTCAAAGATGACAGTAATGGCCATGTGTATGTGTGTGACCCACAACAT
CATGGTCATTAAAGTACATTGGCCAGAGACCACACTGAAATAACAACAAT
TACATTCTCATCATCTTATTTTGACAGTGAAAATGAAGAAGACAGTTCTT
CCATTGATCATCTGTCTCTGAATCAGGTAAGCAAATGACTGTAATTTCTCA
TGGGACTGCTATTCTTACACAGTGGTTTTCTTCATCCAAAGAGAACAGCAA
TGACTTGAATCTTAAATACTTTTGTTTTACCCCTACTAGAGGTCCAGAGA
CCTGTCTTTTATTATAAGTGAGACCAGCTGCCCTCTCTAAACTAATAGTTG
ATGTGCATTGGCTTCTCCAGAACAGAGCAGAACTATCCCAAATCCCTGA
GAACTGGAGTCTCCTGGGGCAGGCTTCATCAGGATGTAGTTATGCCATC
CTGAGAAAGGCCCCGAGGCCGCTTCACCAGGTGTCTGTCTCCTAATGTG
ATGTGTGTGTGTGTCTCTCTGACACCAGCATCAGAGGTTAGAGAAAGT
CTCCAAACATGAAGCTGAGAGAGAGGAAGCAAGCCAGTTGAAAGTGAGAA
GTCTACAGCCACTCATCAATCTGTGTTATTGTGTTTGGAGACCACAAATA
BACACTATAAGTACTGCCTAGTATGTCTTCAGTACTGGCTTTAAAGCTG
TCCCCAAAGGAGTATTTCTAAAATATTTTGAGCATTGTTAAGCAGATTTT
TAACCTCCTGAGAGGGAATAATTGGAAAGCTACCACTCACTACAATCAT
TGTTAACCTATTTAGTTACAACATCTCATTTTGTAGCATGCAATAAATG
AAAAATCTTCTTAAAAAATCATCTTTTATCTTGAAGGAGGAAGGAAG
GTGAGACAAAAGGGGAGAGAGGGAGGGAGCCATGAAACACCAGTTACC
TAAGACCAGAATGGAGATCTTCTCACTACCTCTGTTGAATACAGCACCT
ACTGAAAGAACTTTTATTCCCTGACCATGAACAGCCTCTCAGCTTCTGTT
TTCCTTCTCACAGAAATCCTTCTATCATGTAAGNTATGGCCACTCCAT
GAAGGCTGCATGGATCAATCTGTGTCTCTGAGTATCTCTGAAACCTCTAA
AACATCCAAGCTTACCTTCAAGGAGAGCATGGTGGTAGTAGCAACCAACG
GGAAGGTTCTGAAGAAGAGACGGTTGAGTTAAGCCAATCCATCACTGAT
GATGACCTGGAGGCCATCGCCAATGACTCAGAGGAAGGTAAGGGTCAAG
CACAATAATATCTTTTCTTTTACAGTTTAAAGCAAGTAGGACAGTAGAAT
TTAGGGGAAAATTAACCGTGGAGTCAGAATAACAAGAAGACAACCAAGCA
TTAGTCTGGTAACATATACAGAGGAAAATTAATTTTATCTTCTCCAGGA
GGGAGAAATGAGCAGTGGCCTGAATCGAGAATACTTGCTCACAGCCATTA
TTTCTTAGCCATATTGTAAAGGTCGTGTGACTTTTAGCCTTTCAGGAGAA
AGCAGTAATAAGACCATTACGAGCTATGTTCTCTCATACTAATATGC
CTCCTTCTCATGTTACATAATCTTTTCTGTGATTGAGTTTCTCTACTGT
AAAATGGAGATAATCAGAATCCCCACTCATTGGATTGTTGTAAAGATTA
AGAGTCTCAGGCTTTACAGACTGAGCTAGCTGGGCCCTCTGACTGTTAT
AAAGATTAAATGAGTCAACATCCCCTAACCTCTGAGTGAATAATGTCT

FIG. 3 (27 of 52)

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GGTACAAAGTAAGCACC...AATAAATGTTAGCTATTACTATCATTATTAA
ATTATTTTATTTTTTTTTTTTGGAGATGGAGTCTCACTCTGTTGCCCAGGC
TGGAGTGCAGTGGCGCAATCTTGGCTCACTGCAAGCTCTGCCTCCTGGGT
TCAGGCCATTTTCTGCTCAGCCTCCCGAGTAGCTGGGACAACAGGCAT
GTGCCACCATGCCAGCTAATTTTTTTGTATTTTAGTAGAGATGGGGTT
TCACTGTGTAGCCAGGATGGTCTCTATTTCTGATCTCATGATCCGCCT
GCCTTSGCCTCCCAAAGTGCTGGGATTACAGGCGTGAGCCACCGCGCCCG
GCTATTATTATTATTACTACTACTACTACTACCTATATGAATACTACCA
GCAATACTAATTTATTAATGACTGGATTATGTCTAAACCTCACAAGAATC
CTACCTTCTCATTTTACATAAAAGGAACTAAGCTCATTGAGATAGGTAA
ACTGCCCAATGGCATAACATCTGTAAGTGGGAGAGCCTCAAATCTAATTCA
GTTCTACCTGAGTAAAAAATCATGGTTTTCTCCTCCATCCCTTTACTGTA
CAAGCCTCCACATGAACATAAAACCAATATTCCTGTTTTTAAGATAATA
CCTAAGCAATAACGCATGTTTACCTAGAAGGTTTTAAATGTAACACAAT
ATAAGAAAAATAAATCACTCATATCGTCAGTGAGAGTTTACTACTGCCA
GCCTATGGTATGTTTCTTAAATCTTTGCTATACACATACCTACATGT
GAACAAATATGTCTAACATCAAGACCACACTATTTACAACCTTATATCCA
GCTTTTCTGACTTAGCAATGTATTGATGACATTATGCATGCTTAGACCTC
C

>Contig34

GTATTCTATTCTCGGTTATAACACAATCACAGTGATTGTGCATATCTTTC
CAGGATTTGTTAATTTCACTTCTTCAAGCTGTTTCCCCCTTGTGGCTGGA
ACTGATTTTCTATCTTCTGGGAGAATCTTCAAGCAAGCCAACTCAGGATTT
GTTGGGTGCATTTTGTCAAGTCTAGGACCCAGGCTCTGGGTGACTGATT
CCTCTAATTACCGAGCAATGTAAATGAGGAAGTCTGATTGTGTAAAGGT
GTTAAACTTTTGTGTGACGGCAAACTTTAATACCATGAATAGAGATTCC
AGAATTTTCCAACCTTCTAACGGGATTCCTTCTACTCCCTGACATTAGAAT
GTTAGAAAACTACCACAAAACATCTGTGAGGCTATCCTACAAGGCCCGT
TTTTCAAAATAGGTTTTTACAAGGATTGCTATTTGGGATGATAGTTTCAG
AAAGGCGCTATCAAAGTTAATTGATGATGTGTGCAAGCTGAAAGTTATAT
GTTAGAACTAGCAGTGATTTCAAAAATATCCCTTTTAGGCTTTTTTGCTAA
TATATCTGCTCATTTTCAAAGTTCCCAATATTATAAACTTTTTAAAGCA
GAAAGAAGAACCCTCCATTTCTGCTGGCCCCCTTCCCTGTTCAACTAAAAA
GTATTTTCCCAGGCAATGCTATCCCAGGACTCACACTCCATCCATCCATC
ACCTACCATAAGTTCTTTGAAGGGCTCATTCTGAGCGCTTCTTGAGTGCC
TGGGATCTGTTATTTCTCTCCATTTCTGCTGCTGCATGGTAGTCCAAGTC
CTCCTCCCTTTTCCCTAGGCCATTTGAATCATCTGCTAATTGGTTTTCC
TGATTCCCACGGAACTTCTCCATCCCTTCTCACATATCAGCCACAGA
AGTATCTCCAAAAAGCAAATCTGGTGACATGAAGCCCTTGACAAAAACC
ATTCATTACTGGTTCCACACCTCCTTTGTGGATAAGTTCAAGCTCCTGAG
TGTGGCAAGCAGGGCCACCTGGAATCCCTGCCCCCTCTCTCTATCCCA
CGCATCAATCTTCTGTCTATTTGCAGTTCCTTGAATGTGATATCTTT
CTAGTCTCTGTGCTTTTGCTAACCTGTTCTTCTGACTGGAACTCCTT
CTCCTCCTTGTAGTTTGGCTAATTTCTAGTCTTCAAGACTCAGCTCATG
CTTCAACCCCTCTATAACAAGTCCTTTCCCAAGCTGGGTGGTGGATGCTC
CTCTGTGCTGTGTGAGTCTTGAACATCCTCAGCAAACCTCAGCTTTGTTT
GCTTGTCTCCCTTGCTGTCAATGCACCTGATTGAGGCTGGCATATACTG
TTCACCTCCATGACTGGCTCATGGTGGTGTCCGTGAATATCATCCACCC
AAACGGATGAGAGCTACCATGCCATCACTTGTGACTTCCATCTGGAGCTA
ACCTCCCCGACAGGAAAGCGTTTCCCTAGGAAAGAATATCTTTGGGTTA
AATAGAAGTAGAGACTCACCAGAAGCACTATGTCCAGCTCAGAATGAACT
GCTCAGTAAGCAGCCTTGTCAATGAGGAGGAGCAGGCCAGCCCCAGAGG
CCTCAAAGTGGGAGAGTAGAGAAGCGCAGTTCTGCCACAAAGGCACAGT
GGACACCTTGCTCCCTGGCTGGCTGGAAGCAGATGGTGTCCACCTGCTT
CCATGGGAATTCTGCACCTTTAATAAAGTTTTATGGGACAGGAAGGTGAC
TGGCATTGACATTGTAACGAGGAATGGGTGGTGCCACCTTTGCTGTGTCT
TACCAGAAATACCTGTGGCAGGTAAATTTCTAGAGAGACCCTCCCATTTC
TCCCATATAGCAATTTTGAATGTTTCTGAGGGCTTTCAAATTCATCT
GGGAACATAGGAGTTCCAGAAAGATGAAATCAAAGGTGATGGTATGCCAA
AGAAAGTAGCTTTTAGAATGACTTACATTAGCCATTTCATCCATTGAGCAC

FIG. 3 (28 of 52)

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ACCAGGCATTGAGTTGAGGGGTGTGTGTGTGTGTGTGCGCGCGCGCGCGCG
CGTGCAATGAGTGCAATGCGCGCGCGGTGTACATAGGGGAAGGGAAACAAAAC
AAAAGTACACAAGACATGATAGTTGCTCCTCAAGGAGTTTTTGCAAATGTT
CACAAATTAAGAGAATATGCTGTGCTGTGGCTGGTGTATAAACCAACTGC
TAGGGAGAGGCCTTCCACACACACTTGGGGCAAATGCGACCTCTAGGACT
GCCAGTGGAAATCTGGGCATGCTGTTTGTGGTCGATAAACCCCTGGTCCCTT
GATCAGGGACCTATGTTTACTTTTCTCTCCCTGGAAGTCTTCATTAGTG
GGCATCCAGAAGGTCTTGACAGGGCAGAGGGAGGCACAAAGACAAAGAGT
TTGAAACCAAGCCTGGACAACAAAATGAGTTTCTATCTTTACAAAAAAAT
TTTTAAAAAATTAGCCAGGTAGGATTGCATGTGCCTGTAGTCCCAGCTAT
TCAGGAAGCTGAGGCAGGAGGATTCCCTGAGACCAGGAATTTTGAGGCTG
CAGTGAGCTATTAAGTTGGCGCAAAGTAATCGTGGTTTTTATCAATTA
AGTAATGGCAAAACTTTTAAATGACAAAAACCGTGATTACTTTTGCACCAA
TTTAATATGATTGCACGACTGCACTGTGCTCCAGCCTGGGCAACAGAGTG
GGACCCTGTCAAAAAATAATAATAATAATAATAATGTAAACATGTAAAAAA
ACCCCAAAAAACAAAAAAATGGGTGTTGAGACCCCTGAATTTAGGAAATA
TAGGAAGGAGTGTGATTCTGTGTGTGTCATGTCATGGGTGTGCACCCCTCAGT
GCCTGGGTGGCTTACCCTGGGCTAGTTCAAGTGCGCAAATGGTTTTCTCC
AGCTGGGCTACCACCATCTTCCCCAGGGCCTGTCCATGTATTTGGTGGC
AAGATACCTATGGAAGTAGAGTCCCTCCTCAGAGGAAAGGCTCCTCCCAT
TCTCTGGCTTTTCAAGTAGTAGTCCATGACTTCAACAGGTCCCCACTGCAA
TGTTATGGGTAGTTTAGGTGGGGTCTCCTCTGAGAGCCTCCCATAGCCC
AAAAGGCCCTGTCTAGCTGGCACTGCATCTCCCTCTTCCAGCTCTCAG
CCTTTCTCTTTGCTCATCCCACTCCGCACAGGCTTTCTGCTGATCTTG
GATGTGTCAATCCTGCCCCAAGGGATGCAAGGCAATTTGTCTTTTATT
ATTAAGATCTCTCCTGAGGCCACGTGTGGTGGCTCACACCTGTAGTCCTA
GAACCTTTGGTAGGCCAAGGTAGGAGAATTGCTTGAGCTCAGGAGTTCCAG
GCTGTAGTGAACCATGATTGCACCATTGCAATTCAGCCTGTGTGACACAG
CGAGACCCTGTCTTTTTCTTTTTTTTTTGTAGACAGGGTCTCGCTCTGT
CATCCAGGCTAGAGTGACGCGGTGTTTTCTGCTCACTGCAGCCTCAACC
TGCACATTTTTGTAGAGACGGTGTCTTGCTATGTTGCCAGAGTGGCCT
CAAACCTCTGGGCTCAAGAGATCTTTCCACCTCAGCCTTCCAAAGTGCTG
GGACTACAGGCGTGAGCTACCGCGCCCAACAAAGACCCTGTCTTAAAAAG
AAAACAAAAATAAACAACTCCCTCAAGTCTTTTTTTTTTTTTTGTAGACGG
AGTCTCGCTCTGTGCGCCAGGCTGGAGGGCAGTGGCGCAATCTTGGCTCA
CTGCAAGCTCTGCCTCCCGGGTTCAGGCCATTCTCTTGCCTCAGCCTCCC
GAGTAGCTGGGACTACAGGTGCCCCGCCACCGCCTGGCTAATATTTGT
ATTTTTAGTAGAGATGGGGTTTCACTGCGTTAGCCAGGATGGTCTTGATC
TCTCAGCTTTGTATCGCCCGCCTCGGCCTCCAAAGTGCTGGGATTAC
AGGCATGAGCCACCGCGCCAGCCAGACCTCTTGAGTCTTAAACTCCTCT
GTAGTTCCAGCCACCCTTTAGCACATGACTCTGTTAATTTTGTCTCACT
GTCTGAAATCATCTCCTGTCCACTCTTGACTGACAGGTCTCTGCACTAGC
CCACTGCTTAATCAGAGTAGGTCCCTGTCAACTTATTCATATTGTGTCCC
CATGCCAGTGTGGATGATTAAAAATTGTTGAGTGGAGGCTGATCAGATGAG
CCATCTCCTTCCAAGTCTCACTTGCTGGCTCCTGTCTTAGTTTGTAGTCC
CCATTCTTCAAAGAACGTGAGCCCTGGAAGTATTTTAGTCATTTAGTTC
AGTGCCTTTGGATGGGAGGATCACATCCCTGGGTCCCGTCTCTCAGACTG
TTTTGCTCTAGCTGACTAGGCAGGATTCCCTGCTTCTCTCACTTCGGCA
TGGGACTTCCTTCTGAAATTGCTGCTCAGTCAAGAGAATGACCTTCCCCA
ACATAATCCTACTCCACAGGGACTTAAAGGTGTGTCAGAGATCTCTGCT
CATCTTTCTGGCCAGGTGCCAACGTCAAGTTTATAGCCAAGGGACAAGACT
AGTTAGCAGATCAGGCAGGTCTTAGACCCAGCGTAAGTGCCAGACTTCT
AGCTGCAGTTGTTCTGCCCCACTGGGCGTTCAAGTGGAGAGAGGGCAT
GGCACTACACTGAGCTCTCGGCGAAACCCAGGACTCTGAAATCTCGGTGT
CAGCCACAGGCCACTCTTTTCAAGCAGGACTTCAGTCAGTCTCTGCTACTAG
GCTGTGAGCAGCATGGTAGGCTTTACCCC

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AAGGAGTGTGCTTGTGATAGCATGTGTGANGGGACGAGGAGTAAATAAT
TTCTGCCTTCAAGAAATTGCAAACTAGTAATGGAGATAAAATCAACAGAG
GAACAATTAGAGTATAAGGTAAAAATCTAAGGGCCATAAGAGAGGAGAAGA

FIG. 3 (29 of 52)

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AGTATGGGAGTTTCAGAGGAGGGGGTAAATGAGGGGAGTAGGTGGGTAGA
AAAGGTTAAAGTAAATAATGATGGGAAGGAAGACAAAAGACGACAGGG
GTGCCAAAGGACTCTTAACCTCATCTGAACGGAGTTGCCCTGTTTTGCTC
TCTGATGCTCATGTATCTATCCTTAGAGACAGCTTGGCGGGCAATGTAGA
GCGTAGGGGCTGACATAGGGGGTTGGAGTCCACCTCCGTGACTTCTAGC
AAATTAGCAAACCTTTGCTGCTGCTAAGCCTATAAGGCGGACAGAAATGCC
ATCTTTAAAGCTTGTTATGTAAAGTGCCTAGGACCTCGTAGGCATCAACA
GGAATAATGGATGAAACAAACAACGGTGCCTATCTTGGAGAAAGTGGCA
TCTGAGCAGGAGTATTTTGAAGGTAGGAAAGGGCTCCAAGCACATCTAA
GAGATTAGGGAACGAGAAGCCTTAGCCCTGGGTGCAGATTTAACCAATC
AACTTCTAACACCGCAGGCTGAGAGGTGTGGAGTGAGAGCCCCGCCAGA
GGCAGGAGACCCGGGCTTCGGCCAGACCCCGCTCCTGGTACAGAGGACC
ACGCCCCGGCTCTGCCTGGAGCCAAATGTGGATCAAAACAGCGCGCAGCTT
CCCCTGCTGGTGAAGAACCCGAGCAAGGGGCTCAGTTTCTTTATCCGGA
ACGTGGTGACAATGACATCTCTTTGCAAGGCTGCTGCAGGGCTTTCTGGA
AATACGCCCCGTGAGGTATCTGGGCTGCGCACAGCCTCCCCCGCCAGGA
CCCAGACGCTACTCTGGGGTCCCCTCTGCGCTCCCGGATGGAAGACGC
CCAGGGGAAACTTAGCGAGCGGACGGGACCGGACCTCCCGCGGGACGAA
CTCACTCGGTGGCCTCCTACTTCCCCGGCGGTGTCCAACGCCTGAGAAT
AACGGGAACAGCGGTCTGACTCACCAGCAGCGGCAGCAGCGGTAGGCCCG
GGCCCCACCATGACTCTTCAGTGACAGTTTTTCTTCAAACGCGCGCCTG
TAGCCAGGACCGCGCTGCCGCGCTCCACGCGTCTCATTGGCTCCTGCG
GGTTTGAAGCTCGCTAGTCTGACACGGGAGGGCGGGACAACAGGCAAT
AGGCTCTTTGCGGTGGCTCTGGCCTTGAGAACCCGACCTTGGGGCCCTT
TGATTGGAAGAACGTGCAGCGCACCTCGGCATTGAGGGCGGCTTCTCGG
GGCGGGCGCGCGCCGCTCTGAGTGCCTGTGAGTGCCTCCGAGTG
GGCGTGGGACCTCCGTGGGGGCTCAGCCGGGCTGGTGGTTGGGGGGCG
GTTACGCTGAATCCAGCTGGGGTTGGCGCGCGGGAGTCCCTGGGCGGAG
AGACAGGGCGGTCTCCAGGATGCTGGGGCGCTACCTGATTCTGTCTT
TTCAAAGTCTCAGACTCACAGGAGCTGTGAAAAAATAATATTATAAAGAG
GACATATGGGTCTTATGCATCTAAAGGCTCCTAGTTCTTAGTACTGCAGG
GTGGCTCGTTTAAATGTGGTAAATATGCATAACATCACATATAACATTT
TAACCATTTTAAAGTGTTAAATTTTCAAAGTGTGCAGTTTAGTGGTAT
TAAGTACCCTCACATTGTGGCACAGCCACCCTACTGTCTTTCCAGAAC
TTTTTCATCTTCCCAAATGAAACCCCTGTACCCGTCACTAACTCCGCACTC
CTCCCTCCCCCAGCCCCAGGCAATCACCATTCTAGTTTCTGTCTCTATGG
ATTTGACAACGTAGGTGCCATATAAGTAGAATCATGCAGTATTTGTTCT
GTGACTGGCTTGTTCCTTAGCATAAAGTATTCAAGGTTTATCCATGTG
TAGCATGTGTGAGAAATTTCTTTCTTTTAAAGGGGAATAGCATTTCTGT
GTGTGGAGATGCCACATTTGCTTCTTGGTCCATCCCTCTCCGGACACTT
GAGTTGCTTCCACTTTTGGCTATTGTGAATAATAATGAACATGAATG
CACAAATAACTCTTTGAGACTCTCTTTTCACTTTTGGGTATATACCA
CGAAGTGGTATTGTTGGATCAAACGGCAATCTATTTTAAATTTTTTGGAG
AAACTGCCCTTACTCCTCTCACGGTGATCTCTTGTCAAGGTATATTTTCG
ATTTACCTGATCAGCTGACTATAAGGCCATAAGGCTAACGGAGAAACGC
AGGCCTAGTTTCTCCTAGTTACTAGGAGATCGCAGGCCTCGTTGTCTGA
ATCCCTAGACACACTTCACTCCCTTGTTTTAAATCCTAAATTTTTTTCT
TTTGAAGTTTGTCTGTTTCACTATTCTCCAGTTTCTTAAAGAGGTCTG
GAAAAATGCTTTTGGCTCCTTGTGTATGAAGGTTCTCTTCCATGGATGCT
GGAGAAGTCGTGTGTGGAGGGGAGTCATATCTGGGCACCTGTTGGCCAG
GTTGAGCTTACCAGTTGGGTACTCAGCAGGGCATGAAGCCACTGCAGCAG
CCCTTCTCTTTAGCCGTAATAGGGAGTTTGAAGAGAGCCAGGGTTTCT
GGATTTATGCATTTTGATATTTTCAATAGTGTATTAAATGTTTAAATAG
GAAACTGATCATTATTTTGTAAATGACTGAGAAAGGACTCCTTCAAC
AACAGTTTTCAGAAAAGTGAAGGCGGTTTTGTTTTGGTCTTTGTAGAATCT
AGGTGGTTGAATGCATGTGAGTTGTAGAAGTCACCTTGCCTGATATCCCA
CGCAGTGCTGGAGTATTCACAGACCCCATGTAGGTACTGCACCTTTGCA
GGTATACTGCTGGTGTGGTGGAGCTGCCTTACCTGTCTGTTATTGGAGA
CCCCTGCTTATTAGGAACTTAAATGAACTCAAATGAGCTTCTTGCTT
ACTGGTCTAGTCTTTTGGAGCAACATAGGCCAGTTCTGCCTCGTTTTTT

FIG. 3 (30 of 52)

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TCCATCCTTTGGGTATTTGACGGTCTATTTTGTAGGACACAAAATGTGGG
AAAAATAGCTAGGCAGGTTTAAAAATTTCTCAACTCTACCAAGCATGGTGGC
TTATGTTCTTAATCAATCCCAGCACTTTGTGAAGCTGAGGCAAGAGGATT
GCTTGAGCCTAGGAGTTTGAGACCAGACTGGGCAACATAGCAAGACCTCG
TTCTTTAAAAAATAAATAAATTACAAAAATTAACCAGGCATGGTGGCA
CACACCTGTAGTCCCTTCTACTCAGGAGGCTGAGGTGGGAGGATCACTTG
AGCCCAAAAGTTGAAGGATGCACTGCACTGTGGTTCATGCCACCGCACTCC
AGCATGGGAGGCAGAGCAAGACCCTGTCTCCAAATAAATACATAAATTAA
ATTCTTAACCTCATTTCATCAAAAGTATCCACTGTAGCTTTCCATCATCCTGG
TGTTGTTTTTTTTTAGAAGGATCTGGCTCCATTGCCCGGCTAGAGTGCAGT
GGCATGATCTCAGCTCACTGCAGCCCCCACCCTCTCTGGCTTAAGCGATCA
CCCACTTCAGTCACCCATCTGGGTAATTTTTGTATTTTTGTAGAGATGG
GGTTTTGCCATGTTGCCCCAGGTGGTCTTGAACCTCTGGCTCAAGCGAT
CCATCTGCCTCCATCTCTAAAGTGTGGGATTACAGGTGTGAGCCACCA
CACCAGGACAATCCTGGTGGCTTTTAACGGTTTTCCATTGCTCTCAGGCT
AATGACCTATAAGCCCCCTGCGGGCTTGGCCTTTTACTCCCTGAGCATTAG
CCACCTCCCTTAGCCTTAGCCCACTACTCTCCCTTGCTCAGTGTTAT
CCAGACACTTTGTTTTTTCTTTCCATACTCTCTCTGTCTGGGAATCCA
ACCTTTCTTTCTCATTCTCTAGTTGATTATTATTATTTTTACTCTAGCA
GCCTTATTGAGATATTTACATACCGTACGATTCTCCCACTTACAGTGTAC
AATTCAATTTTCTAACATTTTCATCACCCCTAAAGAAACCTATACTCA
TTAGCAGTCACTCCCCATTCTCCCCCTCTCTCAGCCCTAGAAACCATGA
ATCTACTATCCATCTCTATAGATTGCTCTCTGGACATTTTCATATGTATG
AAATTATGCAATTTGTGGTCTCTGATGGGCTTCTTTTGTACCAAAATAT
CATGGGTTTGATCTAGGTCTCTGCTGCTCGCTGCACAGAAAGCCAGCCACT
GAGATGACAAGTATTGCCAAGGAAGAAGGCTTTAGTCAGGTGCTGCAGCT
GAGGAGATGGGGCTCAATCTCAAATCCATCTCGCTGACCTAAAACAGG
GGTTTGGATAGCAGGGAAGAAATGTAACAATGCGTAAGAAAACAGGAACC
AGGGAGGGGCAAGGAAGCAATCCTGATGAATGAGTGGTCCAAAGTCTCAT
TGCCTGGATGTGGTATCTGGCGAGTTTCAGTTCTTTGATACTTTTTTTG
AGAGGCCTGAAGTCTTTTCCCCAGGAAGAACTCAAACAAAACAAATACA
AGCTTCCAGCTTTAAGACCAGAAGCGTCAATTTCTATGTTTATCCGAAAG
AACAGTCTATGGGACTATTGGTTAAGTTTCACTTTCACTTAGTATGCTGT
TTTCAAGGTTTATCCACATAGCATGTGTGAGTACTTCATTCTTTTATGAC
TGGGTATTCTATTGTGCGGATATACAATATTTTATTTGCCATTTCATCAGT
TGATGGACATCTAGGTTCTTTCCACTTTTGGCTATTATGAATAATGCTG
TTATGAACCTTTCATGTATAAGTTTTTGTGTAGACATATGTTTTCAACACT
CATGGGTATATACCTAATGAGAGGAATTACTGTGTACATACGATAATTCTA
TCTTTAACCATTGAGGAAGTCCAGACTGTTTTCCAAAGCAGCTGCAGC
ATTTTACATTCTTACCAGCAGTGTATGAAAGTTCCAGTTCTTTTACATCC
TCAACAACACTTGTATTGTCCATCTTTTAAATTACAACCATCCTAGTGG
TTGTGAAATGGTATCATTGTGGTTTTTATTTGTATTTCTTGATGACT
AATGATGTTAAGCATCTTTTTATGTGTTTACTGGCCATTTGTATATCTCT
ATTTCAGAGTCTTTGCCAATTTTTAAATTGGGTGAGTTGTCTTCTTCTTT
TTTTTTGAGATGGAGCCTCACTCTGTTTCCAGCTGGAATACAGTGGTGT
GATCTCAGCTCACTGCAACTTCCACCTCCTGTGTTCAAGTGATTCTGGTG
CCTCAGCCTCCCAAGTAGCTGGGATTACACGCACCTGCCACCATTTCCAG
CTAATTTTTTTCTTTGTATTTTGTAGTAGAGCGGGTTTTCCCATGTTGG
CCAGGCTAGTCTCTTTGTTGACTCTTAACCATCCTTCAGTCTCAGACAAA
ACATCCCTTTCTCAAGGATTGTGATTAGCTTGATTATTTGCTTATCTTTC
TCCCTGCTAGTCTGTAACTGAGGGTAGGCCACTATATTCATTGTTCTTG
GCACCAAAATAGAACTAAATTAATGTCTTTTGAATGAATAGGGCTTTCTC
CTTTTAAAGATCCCTTCAATACAGTAACCACACTATATATAAGTAGCCAC
AAGCCCATTCATAATACTACTAGTNCCTTGCGCCAAACC
>Contig36
GGCTCAGCGTTACTATACTGGTCTCAAACCTCCTGGGCTCAAGCGATCTGC
CCCCCTCGGCTTCCCAAAGTGTGGGATTATAGGCGTGAGCCACGGTGCC
TGGCCTCAAATAACTATTTAAGTGAACAAAACCTAGTATGGCACTAATGA
AAAATGTATAAATCCATAATCGCAGAGGGATTTCAACTTACTTCTTTCGA
TTATGTAAAGGTCAAACAGACAAAAGACAATGACAAAACCTTAATGCAATG

FIG. 3 (31 of 52)

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AACACTTTTGAATTAATGAACATATATTGGATATGTACCCAAGAAATTAGA
GAATACATACTAGTTTTGAGTTTATGCAGAACATTTACAAAAATTTAGTG
GAAGCCTAAATTATAAAAAAGTTGCTGTCACGTAGAATAACACACAAACCC
CTGAGTCCGGAATTCAAAGCCCTCCACACTCTCCTCTACCTTTGCATCTT
TATCCTCCACCACTGCGAGTGCATACTCTGGGCTACTACTCACTGTTCT
TGATTCAAATTCATGTTCTGTCAGCTCAAATCATTCTCTGCTGCTGGAA
TACTACTTTCATACATATTCTGCTATTGAATTCTTGTCTTAGCACCCCAT
CTACTCCAAGACGATGTCCAGTTGGGGTTACTCCCTGTCCCATTTTCTTT
GATTACACTTTTTTTTTTCTACTTCCATTATATTATTGATCACATCTGTGC
CACAGTTTTTGACTTTGTGCTGCTTTTACTCTTTTCTAGACCCTGATAG
CTCCTGAAGGGTTGGGTCAATTTCTTTTTTATTTGCTCATTCTCATGGCA
CAGTGAGTGCTTAATAAATGGCTATTGACTGAAATTAATACTGTATCTAAA
TGGACATATTCCACTTCTGGGCCATTCAATCTTTCTTTCTATTGGAACCA
GGAGATGGGGAACCATAACAAAGGTAAGGTTGTCCATGTGAAAGAACAT
GGAACCTTCCCCTGAGGGCCAAAAAGAGCAGGGAAAGGTGCAAGACAA
AATCTTCCATTTTTTAAACAATGTAAGAATGTGGTCCACCTCATGCTCAGG
TGGGACTTTATCATGACGTTATTTTTGGGGACTTATAGCTGCATCATTTA
CCCCATATACATTTACCTTTAGTGTAAGGAACTGAGGACAGGAATTTGT
TGATGCAGACTCTTGCTAATGAGGCTAACACTTGGAGAATTTTATCATG
CATTCAAGAAGCTTGTGTTTACATTTCTTCAATTAATACTTTAGTTGGTGGT
TTAGCTTTAGTTGTAGGCTTATCAGATATTTGGAGATATCTTCATAAAGC
ATGGCTTTGGTTTGAAGAGTTATTCTGAAGCTACTATTTCTGGCAATA
ATCAAACAGCATGGCCATTTGTTTTGTAAGGCCTTCTAGAAATATGACG
GTAAAATCTACGTGTGGAATAATGCTTATCTCTCTCTCTATAAAATGT
GAATCTAGTTTGTCTTCAAATGAAATCAAGTGATTAAATGTAGTTTTC
TAAGAAGATAAATGGAGCAAAGCACTCTGTGTTTACAGTGTGGAAATC
ACTCATCCCTCATAAACTGTCCCACTGATCCTGACTCACATGAATGAA
TTAAAATAAGAGTTAATAACATCAATTTACATTTTTAAAGACACTTTCCC
ATGTTTTAGACTATTGGTTGGAAAAGCTGGTAGGTGTACAATTTGTGGAG
AGTTGGCTGTTTTGTCTGCTGTTGTTGACGTATTTCAAAGCCATATCT
AATTTTGTGTCAGAAATGGTCTGAATTCTACAAAATGTTGAGTTGTGTAG
TGTGGAGAAGTACGGAGCCATTTACTGAAAGGCTGGGGGAAATGACGAG
ACCCTGAGATAAGGCAGTAGTGGTGCGAACAGAGTGAAGGGAGGTAGTT
GAGATATGTTAGAGTAGAATCAGAATGGACATAGTGAACAACTGGATGC
AGGTGGGGCTGAGGAAGCAAAGTTGAGGATAATTCTGAGACTTCTAGGT
TGATCCACTGAAGTTACATTATTCAACACCACAAGGAACTAGGGGAATG
AGAAGGCATACTGGTTTGTCTTTGGAGTGAAGGGCAGTGATGTAAGAGGA
GTTAATGAGTTAAAGTTTGGATATGCCTGAACCTCAATTTGATATGTGCA
CTTGATATACCTTGGGGTGACCTCCAGGCAATGGTTGAACATGTGTAT
TTCTTAGTAAGTATAGGCATCACAGACTCACATCAGTAAGGAAGCAACA
GCAAACCTTGATTGGACGATATACCTGGAACCTCAGTACCCTATGACTGGAG
CAAGTCTCTGTGAGTGAATGAGGATAAGAAGAATCTTGACCTTGTGGAA
TATGTTGTTAGGAATATATGTGATGAACAACATAGGATACTTCTACAGG
GCTCCACATGTAGTAAGGGCTTTATAAATGCTTGATAAATATTATTGTTG
TAATTTATTTCCAAAGTAAGATGCCACTGGAGGAATCTTTGGAACCCAAA
TTAATAACAAATAGGACTGGATGCAATGGCTCACACCTGTAATCCCAGCA
CTTTGGAAGGCCAAGGCAGGAGGATCTCTTGAGCCCAGAAATCAAGACC
AGCCTGGGTGACACAGGGAGACCTTGATCTATGAAGAATTAATAAATAAT
TAACCAGATGTGGTGGTGACGCCTATAGTCCCTGCTGCTTGAGAGGCTG
AGGTGGGAGGATTGCTTGAGCCCATGAGGTTGAGGCTGCAGTGAGCCATA
ATTGTGCCACCACACTCCAGACTGGGTGACAGAGTGAGACCCTATCTCAA
ATAAATAAATAAATAAATAAATAAATAAAGTACAAACCAGCAAACACTAAT
CCTTTCTAGAGATTATTGAACTCTGGAGGGCAGATCTGAATGGAGCCAGC
AGAGGGACCTATGGAGATCAGCCTGGCCCTGGACAGCACCAGGCAATGGG
GTTGCTAGAGAGGTAATGGGGTTGAACAGGGTTAAGCCATGAGGTCTCA
AGAATCCGTGAAGACTCAGACTAATTTTTTTTTTTTTTGCATGAGGATTAG
GTGTTCTTAGGAATTTCAATGAGAGCAGGGTTAATGAAGGAATGCAGGGT
AGGAGAGCTGAGGGAAGGCATCTGAGAGAGCCTGGCTTATGAATGGCTGC
CTCAGTATGGCTCACCTGCTTTCTTGTATCTACTTAGCAGATGATCCCA
CCCCAGGCCTCCAGGGCCAAGGTCAATTCACATAGTCATGGGCCCTTGA

FIG. 3 (32 of 52)

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GGGCTGGAGCAGTGTAAJGAAGACAGAGTCTTAAGAAATTGCATTAAAL.
BTGATGGGTGCTTGGCAAGTGTCTGTCATCCTATGCCAAGCCTGATCTGAAG
GGGTGCATGCTCATAGGTAGCTGCTGCCAAGATTACAGCAGCTTCTTCA
ATCCAGATCCATGCTCTCCTATATTCAATTTTCCAGGGTTCCTGTCTCT
TCGACAGTGATGAGATGCAGAAATGACTTATTGAGTTATTCTCCTGATAGT
TGCCAACTTTTCCAAATGACAAATGGGGCATGGAGCTTGAGAGTGGAAATG
AGGCCCTAGGGATAGCGTGCCTTAGGAAAACACTCCCAGCCTGATGTAATT
CTGGGGGTACAATGGCATTTCATCATCAAGACTGATGTAAAGGGTGACT
AGCAGTGAGTTGGGGGTGACTCGCACTGGGGCTAGGTTTCTGATTCTGCC
TAATCCAGACAGAGCAGAAGCACTAGTGGGCTGGTAGAGGGCCTCCAGGG
CCTCACTTAATGTCTTGGAAAAACAGCTCCAGATTGTTGGTTCACGTTCT
GAGGACAAGCTTGGGTACTACAGGATAGAGAGAGTGGTGGGAGATGCCGT
GGCTGCCCTTCCAAATGACAAATGGGGCATGGAGCTTGAGAGTGGAAATG
GGGCATCTTGCTTCCCTGCCAGACCTGTAGTTCAGCTGAGGGCATGTG
GAGGCCAAATGGCTTCTTAGAGTGTTACTTTCTTGAACAGCTCTGCTGG
GAGAACTGGAGGAGCTAGCTAGTCACGGTAACTGCAGCAGTCAAAGGATC
GTCCCGGTGGAGGTGGGGTGGAAAGGTAGAGAAAGAGAACATATAGCGTT
TTCCTTGGAGATGTGTGGGCATGTCTAGAGGAAATACCCAATTCTGAG
CCTTGAGCCCTCCAGGAAACCTTGGAAATATTAGGTTAGTCATCCCCAAGG
AAGTCTAAGAAATTCTGGTCTCACCCATCTCCTTTAATTCCCACAATGATC
CTACATGATTAATTAAGGAACCGGGCCAGTAACCTCCAAGCAATGGATGT
GGTGGTGAAGTTTGACCTCATGATGGAGCGGAGGTTGGTTTGAACCTAA
CAATTTAATTTATTGTTTCAAACCTGTTCTCCACTCAGCGTTATTAAAGCA
TACATAATTGACACATAAAAAATTGTATATGTCTACGGTGTACAATGTGAT
GTTTTCGATCTATGTATACATTGTGAAATGATTACAACAAGCTAAATAACA
TACCCATTTCATCGTGTTCAAAGGAATTAAGCTCAAGCACAAAAGAGAGG
TGCTGTTGAAGAGTAGGGCTGCTCTATCTAAGTAGTATGTCTGGGGTGT
CCTGGATCAGGCTCCTTTGTGCTAGTAATAAACAGCCCTTCTGGGGCT
GCTCCACTTTCCCCACATTTTCTTCTGGAGCCTCCCTAAGAAATTAGGACA
TGGCCACTTTCTCTGCATAGGCTTCTTACTTCAACAAGGACAGGGCTTGT
GCTGCCCCATGCCACTTGAGTGTCCCTACAGCACAGAGCTGAGTGACAC
TGGCTGAGTGAGGAAATCCCCAGATTAATCTTGGTTCTAAGCATCATGG
CTGTATTTACACGTATATGAATTAACAATTACAGCATAGTCAATAAGG
ATTTTGTGTCTACAACCTGGAATCCAGATTATGCAAATTGGATAGTATAA
TATTGAATTCCTAGGACTTTTATTAGTTTTAAAAAATTATACAAGCTT
AGAGTAAGAAATTAAACAGTGCAAAAGAAATCACTGTGAAAAGTAAATG
CTCTGTCTCTGCTGAGAGACAGATATTGCAGCCAGATACTACTGGGGTC
AATAGTTTTCTTTAAGCATGCCATTTTGATGGTTTATGGGACTTACAGCT
CAAGAAGCTTGACACTAGGGTTGATCTCAGAAAATCATTGTTGCAGGTAT
TAGATATGACCGTCTCATAAAGATACACACACAGACACAGCGATTGGAGA
TATTCACTGGGGCTTATGGGCTGCTTGCTCTTCTGCTCTGTGCCCTAAGT
TGGGCTCAGAGTAGCCTGGCATCGGCTGTGGGGAGAATGCTGGCATGGGG
TTAGCAGGAGCCCACTTAACATGTCTAAGCCACCTGGAAGAGTCTTCA
AGGAGACCAGACTCCAGAGGCCCTAAGGAAGGAAGGACTTTTGCCCGTTT
TTAGGTATTCTAGTCCAGAGTTTATGGGAGGAATGGTTTGGCTTTGGGTC
GTGTGCCCCCTTACCGAGTGGGATGGGATGTGCCCATGAGCTGTTGAGCT
GGCTCTTGGAGAAGACAGCAAAAGCGGAATAAGAGGTCAGGAAGCTGTG
TGGTTGTAGGAAATCCAGCAGAGGGCTGGGGGTCAAAAGTGGTCATGG
TAGTGACGGTGGAGGCTGAGGTGGTAGAAAAATCAGAGGACAAACCCCATG
GGCTGCTGGTGATCTGACCGAGCTCCTATGCTCTCCTGGTTCATTTTAGG
CTCTGTAGCAGCAGATGATTGGCTGGTGTGAGAGCAGTGACCTGCCATA
TCAGGCAATCCAAGACAAGTCCAAGCTACGCTGGGAGGAAACCTGAAGGC
AGCAGCAGGTAGACTGGCTGAAGACAGACAGGCAGGCAACTTGTCAATCA
GATTTGTGTTTTTAAGGACTTTTAACTGGGGAGCCCTCCGGGACAGATCA
GATGAGAGTGAATGTGCTCCGCTTAGCC

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GGCCGTTCCCAATTCTGTAAAAGGGAGAGTGGTTTTATTTATTTTAAAC
ATAGTCAAGCTGCTAAAGTATATGATATGTATAGATAGAGTATAATTAA
TACTTTCAACTACAGACAAAATCAGGAGAATGGAATTAACAAATTTA
CAATGGGTAAATGGCAGCATTGGGTTGCGCCACCCACGAGAAGGCAGAC

ACCAAGATTCTAAGATC...ACGTGGCCAGCACTTCAGACTTCAAATAGAA...
TTCGTGATTATGCATTATTTTTCTCGGAAAGTTTTCACTTCACTATATGC
TACTTGACACTTGCTTTCTTAAGACATCCCTCTATTTTTGAGATGACTAA
CTCAGCAATTCAATTTCTCTCACGCATAAGCTGTCACTCAACCCAAACCCA
CCAAGCCTGCATTCTACCCTCAATAAGGTCTTGGTGTGTAAACTGACCCA
CTTACCTTAGTTCTTAGCCCTCTCTTGACCAGACATGACTCTTTCATAA
GCTAGACCTATAAAGTCAGGGCTCTTAAGTAGCTGATCTCTGATAGTGCC
AAGTGTCCSSCACTGTTACATTTTCCACTCCAGCTTCTAACAGGTGATA
GACTGCTTTTTGGGGGTAGGGGCACCAAAACATATAGACCTCATGTTTG
ATGTAGACACTCCAGTTTCTTTAAATTACAACATACATATTAATAATGACT
TCCAAGTGTACATTTTCAGTCCAGATCTCTCCCTGGATCCCCAACTTTGT
AAAACCCACCGCTTAGTTGATATCTTTTGATGTCTGACAGGCATTTCAA
TTTAATACTGTCAAAACAAAGTTATTGATTTTCATCTCTGCATCTGTTA
CAAATTTTTCTTACTTTGGTAAATAGCACCCAGGCTGTGTCACTGCCAA
GAATTTTCCACAGCTCTTGAATAAAATTCAAATATTTTCCAAGGCAGA
AAGGCACAGTGTAAATCTGGCTCCTGCCTACCTCTCCAACCTCGTATCACA
CTAGTCTCCCTGTCACTACCCCTCCAGGAGCTCAGGTATCCTTAAAGT
TTCTTTTCTTTTTTTTTTTTTTTTTTTTTTTTTTGAACAGTTTTGCTCTGTT
GCCCAGGCTGGAGTGAAGTGGCATGATCTCAGGTCACTGCAACCTCCGCC
TCCCTGGGTCAAGTGATTCTTGTGCCTCAGCCTCCCAAGTAGCTGCAATT
ACAGGCGCGTGCCACACACCCGGCTAATTTTTGTATTTTTAGTAGAGAT
GGGGTTTTCAATGTTGGCTAAACCGGTCTCAAACCTCTGACCTCAAGTG
ATCTGACCACTTCAGCCTCCCAAGGTGCTGGGATTACAGGCGTGAACCAT
TGTAACCTGCTCCTTGAAGTTTCTTGATCCAGACTCATTCTGCCTTAA
GGTCTTGCACTCTCAGTCTCCCTCAAATGACACCTCCATGAAGACGCA
ATTACCTGTAATTACCGTGTCTTATTTAGTCAATGTGTTGGTTTTCTGTC
TCCTCCACTACAGTGTAAAGCTCTATGAAGGCAGAAACCTTGGCAGTCCAG
TTCCAGCACAGTGCCTAGCACACATAGGTATTTAATAACACACAGTAAA
ATTCACCTTTTAGTGTGCAATTCTGAGTTTTGACAAATGCATCAAGTCAT
TTAAGTCTGACTATTATCAAGCTATAAGATGGTTGCAACACTATCACTAA
TTCCCTCATGCTCCTTGGTAGTCAGTCTCACCCCTAACGCCCCCTCCTG
GCAATCACTGATCCGTTTTTTGTCTTTATAGTTTTGGTTTTTCCAGAAATG
CCAATAACTAAGTTTTGAATGAATGAATGCTATTAACTCTCATTTCTGAC
TCCAGAGCAACATCCATGCAATATTTATTATTTACGCCCCAAATACTGCC
CCCTCACCTCACTCCCAACCACTACTTGATGATACAAGGTGAGACATT
GGCATGTGCTTCTCCATGTTCTTAGCATTTTCCCTATCTCCTTAGCCTT
CCTTCTAATCATAAACGAAGAGTGAACCTTCCCTTTCTAAAGGCAACTTA
CTCCTAGGACCTCGATGCCATAATTTGTTTCTCTAGTACTTTCTATATA
TACACCAACAATTAGCTCCAGAAAGGTAAAGACTCACTGTGTGCTCATC
ACTGTGTCTCTAGCGCTGGCACACTGCAGGTGCTGAAGAAACACCTAC
AGAATGAGTGAATGAATCTCTCCCTCTCTAGACTCCTTCTCTTTGTAAT
CAAACATGTTCAAGTCAACACAGTCTTATGACCAATCCTCTGTTGTCT
GACCTAGGCTGAGCTCCAGGGCTGGGACCCTGACTTCTTATTACCACC
TCAAGGTCTCTGCACTCACTTCTCTTCTGCTCAGGATTGTTTTCTTCT
TGTCAACAGTCTTTTCTCAGACTTAGGTCTCAGCTCAGACATTGCTGTTG
AAAGTACTTCTACTGATCCTTTTATCTAAAGCAGCCATTCCAGCCCTACT
CTCTTGATCATAGCACCTGAATTAAGTTGTTTACTTACTGTCTCTTCAG
GAGGGCAAGGAGCTTGGTGGTGGTGTTCAGGGCTGTACCAAGCTGTACCT
TGCTTACCCTGCTACACTTTTTAGCAACCATCTAATTTTACATGCTCCC
TTCACCTGTCAGAAATTTCTTATTTTCTACTTCAAGCAGGTATACATAT
GTGCTTCTCTGGGAGGCTCACCCACTTCATGAGACTACATTTGGTCTG
GGTAGAAAGGTACAAAATCCACTGGCTCAGTTTTAATCAATGTATGTTA
ATATTAACCAACCTGAGATCTTGATTTCCACGCCTGGCTAATTTGTATT
TTTAGTAAAAACAGGGTTTTCTCCATGTTGGTCAGGCTGGTCTCGAACTCC
CGACCTCAGGTGATCCGCTCACCTCGGCCTCCCAAAGTGCTGGGACTACA
GGCATGAGCCAGCGTGCCCGGCCTAAGATCTTGATTTCTACCATCTGAAC
TCTGTATTTGAAGTCACTGCTCCTGCTTGAGCTTACTGGCCAAAACCTGG
CCCACTCAGACTCACGGAAGTTTCTGGTCTTCCCTGGTAACTTTCTGA
ACTTAACCACTGGTTTGCTTGACAAGAGATTACCATCTTCTCACTTCCTA
GCTATGTGAACCTACTTATCTGCTCTATTGCTGTTCACTCTAGCACGGCA

FIG. 3 (34 of 52)

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CTTATTGAACGAGTGTCTACATCTGCACCCCTACTTCTTACTCATCCAT
TCTGTTTCAATTTCTTAAAAAGAAAAAAGCTATTGTAAACATACG
ATTACAGAAAATGATTTATAACATGTGTATGTACCACCTAGCCCTGTCAA
STCTTAATATTTGTTATATTTGCTTCAAATCTTTTTCAGACTGTAGTTA
AAAATTACTTAGGAGCCATTATTTATGGCCTATTTCTGACCTAGTCTTC
TTGATGGTCAATTTGCCTAATCATCTTAAGTTGCAAAAGCTTAGAATTAA
AGCAAAGTACCTTCGATCCTCTGCTGTGCTTCTTTTAAATATTTGGGT
TTGTTTGGGTCCCATTTACGGTTGTGACATCAGCTTGAGTTTGGGAGCT
GTCTTGTTCAGAAAATGGTTCTGGGAACAGCCTTTTCAACTTGGAGTC
CAAAGTCTGTGCTTTTGTCTGAAAGCCATTATTGTTATGTTTATTACCAC
TGGTTCCATTTGGTCTTATGCTAGGGGTGCTTGGAAATGGCTGAATTAAT
CTGCCAAGTCAAAATTAGGCCTCTGGCTTACGGCTTTTGACTTTTGTCAG
TACACATGATGTCTGAGGTATACAACTTGGCTGGACTTCTGATCTTGCT
TGATGTTTGGATGTCTGTTGTTATATTCACCCTGAAGCAAACTGGGGTAT
GTTCTGGGTTTGGTGTGCTTCACTCTCTGTTTCAAGTAACAGGGTATGACC
TATCTTAGTTTCATTTGGTCTTTCATATTGACTCCTATTAACCTTTATAT
CTTTGATGTTCTTGACTACTGGTTTCTTTGATGACTGAACCTTACTAAGG
GTCCGAATAAAGTGAGAGGGAACCGTCTTGGAGGTTTTACTCCTGGTCT
TGCAAGATCTGCTCCTCTAGAGAGTTGCTGTGATTTTACTGGGAAAGTCC
TGCTTTGTGTTTCTCCAACAAATTTGTTTATTAACCTATCTTTCAGAACA
GCCTATTAACTGAACCTTTGCCAAGGCTTGGTTAGGAACTAACTGTT
CTTGGTTTGAATTATAAGAGTCAGTCTTTGGCTTACTTCTGGTATATAATT
TAGGATCTGGCTTCTCTCAGGTTCTGTTAAGATATCTAGCAAGTTCTCT
TTGTTTGTGTTTCTTTAGAAAGTTATCCAAGATTCTGTTTCAACATGGAT
ATTATTCTAAAGTCTATACATTTACCATTTCTTGATCTGTTAACTGCT
GCTTTGTAGTTTCAATTGCTCTATATTAAGTGACCCACAGGTTTTCTT
GACAGTCTTCTGTGGTGGACTATCTAGCTTCACTGTTGAAAACCTCTT
GCTGAAAAGCTTAGACTATGGGTTAGAAGAAACACATTTTGAAGTCCGCC
TTTTGCCCAGAAGTTTGGTGGCTCTAAGTTTCAAGTTCTGGGACCTGCA
GTATTAGGTGGTCTGGGCTGGAGTTTAATGCTGATGGACCTTTTAGGTTT
GACAGGCAAAACAACATGGTTGGTAACATCATTTTGGGTCTAATAGTCT
GAAAAACAAGAAAATACATATTAATAAATCCTTAACATATCTTATTGT
TTTTAAATAAATACTGTGTTTAAACATGCTAAAAAATAATCATTTTT
AGAATTTCTCTAAGAAAGTTGAATCCTCAGAAAGTAAAGAAAGACTCAC
TAATAGGTAGTTTGTGTTTTTTTTTTTTTTTTTTTTTGGAGACAGGATC
TTGCTCTGTACCCAGTCTGGTGTGTCAGTGATGCAATCTTGGCTCATTC
AACCTCTGCTCTGGGTTGAAGCAATCTCCACCCCAACCTCGCAAGT
GGCTGGACTACAGGCGCATGTCACTACACCTGGCTACTTTTTGTATTTT
TAGTAAAGTTGGGTTTTACCATATTGGCCAGGTTGGTCTTGAAATCCTG
ACCTCCAGTGTACACGCACCTTGGCCTCCCAAGTGTGGGATAACAGG
TATGAGCCACCACACCTGTCTTAACAGGTAGTTTTTACAACCTGAGTTCC
TATCAGAAGTATATTAGAATCTTTAGCTTGACAGAATTAAGCAGAGATG
CAGTGAATATACAAAACCTTGCTCTTTCAAAAATGAATTTGCCTCAAACAG
TAGTTGTTGAATGCCTATTATATCCTAAGTGCCCTCCAAAGAACCCTGAA
AAAATACATACATAATGAACTTATGTTAGGGTACCTCCCAACAAATCTCT
CCTAGTACTTTGTATAGCCACACTATATGTTTTTAAACCACTGCCTTTG
TAAACATCACAGTATCACTCAAGAACCTCTGTCTCATCCCTGGAGATCAG
TGACAAGGAGATAGGTGGCAGATGATGTGAGGCCTGAGATATGCTGCCAC
AGCTCTCAATAAACATGTAACATCTTAATAGTCATATTTGTAAATCAGC
CAGGACAGGTTTTTAAGGTTAGAGTCTATGTTAATAATAACAAATGTTT
AGTCATGTGATTTAAGTTTGGATAAGAAAGGTAGGACTCGATTACAGAGA
ATTTTGAACCTAGGGAAGGGAGTTTAGAATTCATATGGTAAGTAATTGG
GCAAGCCACTATGAATTCCTGAGCATCTCTCATGAAAGCAATTACTCAGA
AAGGAGAATTTACAGAGATTTATGGAATATGTTTCCAGGGTAAGATATG
GGAATGCTAGAGTTACCACTCTATTTTTGATTTGACAAATATTGTGAAGA
ATCACTACATAAACTTGGCGAGTATGTAAAGGATTTCTAACCAGAACCAT
TTGGCATTTAGGGCAAGAAATGTCTACTCTGGATGATAGCGGTGTGTGT
GGTGTACTAGGAGTGAACAGCGGAGTTGGAGTGGGAGGCAGAGAGAT
GGATGGTATACCCACAATGGCTATATCTGGATTAATCTTTGAGCACCAC
ATTTATATACACCTCGGATCTCTCCATCATTGCTTACTGAAGAGGTGGAG

FIG. 3 (35 of 52)

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GGACGTTGGCATGAAAGCTCCAAATGTGTTTTTTAGTTGCTTTCTTA
ATATTAAAAACGAATTGATATAATCCACAAACCATAAAATTCACCATTTT
AGTAAGTGCACACTTCTGTGGATTTTAGTATAGCCACACTATTATACAGC
AATCACCACCTGTCTAATTCAGAACATATTCATCACCCTAGAAAGAGAC
TTGGGTTTACTTGTGGCAGTCCCTCCCA
>Contig38
GGTCTACATGTGCTCGCAAGATTGGATATTGAAATATCAGCAAGAAATTA
AATGACATAGTAGTCATTATGCCTAAATTATTGTTATTTTTTGATTGAAA
AAAGTTGAATATTTCAAATATCAAGGTAGTAGTGAGATATAATAAGAGA
GAGTCAGTTCTAAGTATAGAATTGCTGATTAGTTAAGCTCTGTTCTCCA
ACATTTTGGGCCACATTGAAGAGACCATGTAGCTGCTTTCAGCCTCGGTTT
CCTCCTTTGCAAAATGGGGATTACACTACCTGCCTCACAGAGATGTAAAC
TTATGACATGTTATCATGATTGCCAGGGCCACCTGTTTTCTTTTAAACA
TTGAAATCACTGTGCTGAAACAGGGATTTCCCTGCCCTTTGTGCAAGCT
CCAGAAACAGGAGTCAGCCTGAGTCCCGCAGCTAAGAACGTGGATTCTGG
TCATTTTCTCATAGCGAACACACTTTCAGAGTCTTCAAGGGAGTACATT
TTCCTATAACTCACCTTAATCTCAGTTGAAGCCTCGTTTCTTATTTTGCA
CTGTGGCCAAAACTAAATCTCATTCTTTTACGTAAACTTCAGCAATTC
AATAATAGTACAGTCATTTTATGTTTCAACTGAACCAAGTCAGGGTTCCA
CTCCTGCCTCCCTTTCTGCTCTGAGGACATCCATGAAGTGGAGGGGCTC
TATGTAGCCTGGAGCTATTGGTGAGGGGCGATGGGTCCGTGGTGGTCTTG
GGAACTCGCGGGCTGTGCTGTGGCTGGTCTGGTCTGGTCTGGTCTGGTCT
GTTCCACGCGGGTTCAGGCTGCAGGACAGTTCGTGCTCTTCTTGTCTAAT
GATCAGCTTTTAGGCTCACGGCCTGTCTCTGCTGAGATATGGAATAGGA
CAGCCTCTGGATCTTCTTAACTCTCCTGGGGCCACAGGGGACTCTGTT
TGTGCTGTGCCCCACATAGGATGATTCTGCCAGACCTTTGCTGCCATTT
CTTGCTGTGCTGCTGTTTTAGTCTCTGGAGGGCTTGCAGTTTCTTGGG
GTCCCTGTGGAAGCAAAGCAAAGTCTCTCCAGCTCAGATGTCTAAACG
TATCTGGGTTTTATCGTCCACCCATCCAGAGCTCAGTCTAGAGGAGGGG
GCAGCCTTCGGGTTCTCTCTCTCTCCAGAGCCTCTCTCTTGCACCAG
GGCAGCCTCTCTCTATCTGTTGGAAGGGCTGTCTGGTTCTTGAATATAG
AGTTGCAGGTTTGGGGGTGTAGGCTGAGGTAAGGCAAACATCACATGG
AATAAAATTACCCTGTGTCAAGGAACAACAGAGCTGGACAGTTTTTAA
ATGTGAAAACCAATTTTATTAGGACTATGGCGAGAGGTGAAGTAAGACC
TCAGTATAGAAGTGGGCTCAATTCGAATGCAGCATGGGCAAATGGGAAT
GTATAGCCTAGGAGCAGGGTGGGAACCTGTGGATGAAGAATTACTAAAAG
GGCATATCAGGGGTGAGGGGCGTCTGGCTACACCCACTAACTACTGTT
GCTGAAGAAAGGCTGGTGACATCACTGGGAATGGTGGGGATGAAGAA
TCCAATCAGATGGATATTGAGGATAAGGGGATCTTGATAAACTGGCTTAG
GAGGGTTTTTGTCTAAACTGGTTTTTCATAGGTAAGTCCACAGACAGGTCT
TGGAGAAAGTTTCAAGGACCTACGGTTTTGTTCCGGGCAGATGCTTTGTCTC
TGTACACTGGCACTGTACCTGGCTTCTCTTAGTCCCTCCCCCTTT
TTTTTTCTGGAGTAGTTTGGGAGACCAGAGGAGCAGGGAGTTAGGGAG
AGTAGTCAGAAAAGGCCAGAGAAAATAAGGAGGTGTCTGTAGGGAAAATC
CTTAAATCCTCTAATTAAATTAATTTAATTTATTTATCTGGGACAAGGTC
TCACTCTGTTGCCAGGCTGAAGTGCAGTGGTGTGATCTCGGCTCACTGC
AGCCTCGACCTCAGGGCTCAAGCAGTTTTGCCACCTCAGCCTCCTGAGTA
GCTGGGGCTCACAGGTGTGCACTACCATGCCCGGTAATTTTGGGTTTT
TTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTGTAGAGATGAGGTTTCGCCATG
TTGCCAGGCTTGGTCTCGAACTCCTAAGTGTCCATCCACGTCCGACCTC
CCAAAGTGCTGAGATTACAGGCATGAGCCACTGTGCCCGGCCTAAATTCT
CCAATTTTAAATGCTTCCCTGTTCCCTGTTCCAGATTTGGGATATTGAC
TGCTGTTAAATCAGCGATTTCTCCCTGTGGAGAGGTAGCCAATAGGAAGC
AACAAGAGTGAGGAGTCTTATATCGAAATAGAGGGTAAGAGAAGAGACA
GATGTTATCTTGGCAGTGATTTAAGAACAGCGAGTCTGTAAGCAAAGCAA
AGCAAGGCTCCAGGTGCTGAGAAACAATGGCTTCTGGGGAAGCGTCTG
TGTTCAAGACCTTAAGTTGGAACATCTCTGAAGATGTTGCCATGAAGG
TTTTCTTCTGAAGTTGAGTCTTTCATCACTAGGTAGGCGTGTGTTGGAGT
CTCTATCAAACAGATCCTGTGTTTATTAGGAAGCTGTGGTTCATAAAGCC
CCATGCTAATTTTGCAGGTAGCAGGGTGGCCCTGGCCTGACCCGGGACA

FIG. 3 (36 of 52)

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SAGTGGCTGTCTCCCTCCCTCAGGCAGGAACTCTCTCTCTGCCACCTAGTCT
CTGCATACCCACATTTCAAGGGGAGCTTCTGGGTGGTGTAGTTTACCAGACT
ATGGTCTGAGGTAGAGTTAAGCAAAACAAACTAAACTGCATAAAGAAAC
AGAAAGAAAATCAGGTGTTTATAAAAAACAATTTGGCATTGTGTTGTGTTT
AGCTCCGTGTGATTTTATTGCTTCCACAAATAGTGCCGATATGCACCAGG
CACTGTTGTAAACTGAAAATATGTTTTTGGATGTGCCAGTCTGTGAGT
ATTAAACGATGGTTGATTTGAAATTTGCTATGATTATATTTCTGGGGGT
AAGATGCAGGATTTCTTTGGGGGGCCTACGATGTGGCATTCTAGAATTCT
CAAAGAATCAACCCTGGTGGGACCAGGAAGAGCTGAGCTGAGGCCTCTCT
GCTCATGTGTACTTACTGGAGATCATGGAGACAGGTGAGCCTGAGTGCAC
GTCTCACCAGGACAGCAGAGGGGGAGGAGGCGGAAAGAGAGCTCTCT
CCATTTCTGAGAAGTTAATGGTAACAATGGCATAACATACTACTTTACAG
TTGAAATTTGAAACCACAGCATTAAAGTGTTCCTCAATGAAATTTGGCAATT
TGGGAGTTTTCTGAGCTGCATTGGATGTGGTTTTGTCATGCTGTTAGGATG
AGCAAGAGATGATGGAGAACATCTTCTTTTGGAGCTTCTCTTGGACGTG
GGTCACTCCCACTCATGGAATTAGAAAGCTTAGACCTAGACTTGAATCTC
ACCTTCTCAAGGTGCTCCCGGGCAAATCACTTAAGATCCATCTTCTCTC
CTCCTGCTCCTTCTCCTCCTTCTGAGTTTTTTTTTTCTTTCCAAAATTC
AAATGACACGGTACTGGTAGAAGAAAAGGTCCAAGTCTGCTTTTACAGCT
CCCCCTCATCCCCAAATGTACTCCGACCCCAAGATGACCATGTTATCATTT
GATTGACATCCTTCTAGTTTCAACTCATTTCTTTGTCATGTATATGCACGT
ACATATACACTATTTTATTTTGGCAGGGGTACCGTTTAGCTGCATTAAT
TTCTTATAAAATAATCTATATTTACTTATGGTTTACGTAAAAACAACATAC
ACATGTAAGTGTATAGCTTGATAAGTCTTCACTGTAAACCAAAAAATAAA
TTGGAAGCCCCCAACCGTCTGAATGGACCCCTCTTCTTGGCCAAGAGC
ATTCCAAAGTTAACCTGAAAAACTAGTTCAGGTGATGGAAGGGAAG
GTTGGACATGCCCCAGTATACCCCTTCTCCCTTTTGGAAATTCAGGAAAAGC
TGACCAGCATTAAACATCAACACAGACCTTATGTCTGATAGGAAACTTTGA
CAATCTATTCCCTCTGAAGCTTGCTACCCGGAGGCTTCATCTACAAGATA
AAACCTTGGTCTCCACAACCGCTTATCATAACCCAGACATTCTTTCTGT
TGAGAATAATTTACCTTGTAACTGGAAGCTCCCTGCTTCAAGTTCCCTC
ACCTTTCCAGATTGAACCAATGTAAACCTTACATGCATTGATTGATGTAT
TATGTCTCCCTAAGATGAATAAAGCAAGCTGTATGTTGACTGCCTTCAG
CACAGGTTGTGAGGACCTCCTGAGGCTGGGTACGGATGCATCCTTAACC
TTGGCAAAATAAACTGTCTAGATTGACTGAGACCTATCTCAGATACTGTT
GGGTTCAAATATATACTTATGAACTAATAACAAAAATCAAGTCATAGAA
TATTTCCATCACTCCTCATCTACCCCCAAATTTCTTATGCGTCTTTGCA
GTCAACCTCCCAACCCATCCCCAGGCAACTGCAGATCTACTTTTGTCTC
TGCACCTTCAACTGACCTTTCTGTGATTTTATATGAATGGAATCATGCG
CTGAGCAGTCTTTTGTGTCTGGCTTCTTTGCTCAGCATAATGTTTTGA
GGTTTGTCCATGTTTTTGTGTTTGTCAATGGTTAATTTCTCTCCATTGCA
GAGTAGTTTTCTATTGTACATGTGTACCACAATTTGTATATCCATTCCAT
TGCTGATGGACATTTGATTTGTTTTCCAGATTTTGGCAATTATGAATAGAG
CTACCATGAACACCCAGGTACAAGTCTTTGTGTGGACTTATGTTTTTCAAT
TCTCTTGAATGGAAGTGTCTATCAATAAGTATATGTTTAACTTTGTAA
GAACTGACAACAAATTATCTGCGATGGTTATGCCATTTGTTTTTCTAC
CAGCAATACACGAGCATTTTCACTTGTCTCCACAACCTTTGCCAAAACCTGTT
TTCTTTAATTTGGACATTTAAGTGGTGTACAGAGGCATCTCATTGTGGTT
CTAGTTTTCTTTGCCCTGATGACCAATGGTGTGAAACATCTTTTATGTG
CTTTTTGACCAATTACATATCCTCTTTTGTGAAGTGTCTGTTCAAATATT
TTTGCCCATTTAAACATTTGGGGGTTTTGTCTTATTATTGTGTTGGGAGA
GTTCCATATTTATTTATTTATTTAGAGATGGAGTCTCACTCTGTTGCCAGG
CTAGAGTGCAGTGGCGTGATCTTGGCTCACTGCAACCTCCACTTCTGGG
TTCAAGCAATTTCTCTGCCTTAGCCTCCTGAGTAGCTGGGATTACAGGCA
TGTGCCACCACACTGGCTAAGTTTTTGTATTTTTAGTAGAGATGGGGTTT
CATCATGTTGGCCAGACTGGTCCGAAATTCCTGACCTCAAGCAATCCACC
TGCCCTGGCCCTCAAAAGTGTGGGATTACAAGCATGAGCCACTGTGCCT
GGCCCATATTTATTTTTATTCTTTATTTTGTATACAAGTTCTTGGTCAG
ATACAATAATACCTGGTCAGATGAGATAATGAGTTGGAAAATGCTTTGCA
AATGGGGGAGAATAATTTAAATGTTATTTATTTATTAAGAGCAGAGGCCC

FIG. 3 (37 of 52)

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TTCTCTGTTGCGGTAC...AAGCCGTTTGCTTCTTCTGCCTTTTATAAA...
AGCAGAGTCGAGCTACACAGGCTGTCTGTGTTGGCTGCTATTAGTTAATC
AGAGAGTCTTTTCTTTCCTTGCCTTGTCTATTCTAATTTGTGACACATAAT
AGCCACAATATGTGTTTTAGTTGTGACACTGGCCTGGGAAACCAAGGGA
TGTTTAGAGTGGATTTCCTTGATTTCGCAATAATTGTGTGTTTTCTGCA
TCTTCTGTTAAACACAAATTCATGGAAGCAAAACATGGAAGCAAAGTACC
CTGGACATCCCCCTTCTTTATGAAATTGATTTCTCTTAAATGTAATGTT
TGCTTGTTCCTTACTTTAAAGCAATTTAAGAGTTTATTGAGAAAGTGA
GCCCTGGAAACATAGATGCATAGAGAGAAAATTCTACCACCCTCAGGTCC
CTATTGCTCTCTCATAAAGTGTAGTTTCAGGGCCTTTTAGAAGTTTCT
TTTCTGCTCTGATTTCATGTTTGTGAGTGTGCTATTTTAAAGTATTTGG
ATTTGGTCTGCAATCCTATGAGAGATGGCAACAGAGTAGGGATCTCAA
GCCTGCAAGTTGTATTAAAGTCCAGCAGGGCCTTGTATTTACAACAGAGGG
CCTTTGAAGACATTCCATATATTATGCTAGGGGAGTGGCCAAGCAAAT
TAATGTGTCCCTATGGTGGGATATTTGGGGTTAATACCTGCCCTTCTCTT
AATTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTTTCTTT
TGAGTCTTGCTTTGTCAACCANGCTGGATTGGAGTGCAGTGGTATGATC
TCAGCTCACTGCAACCTCCACCTCCTGGGTTCAAGCAATTCTCCTGCCTC
AGCCTCCCAAGTAGCTGGGACTATAGGCACACACCACCATGCCTGGCTAG
TTTTTTTTTTTTTTTTGAAACNGAATCTCGCTCTGTGCGCCAGGCGGGA
CTGCGGACTGCAGTGGCGCAATCTCGG

>Cont: 39

CGCTCGCATCCCTCATATCCATGAGTGTCTGTGGGCCCTGCCTCTGAAA
TAAATCCTGCCTTTGTCTCCAGTTCACCTCCAGCCACCCATCCTGGGGCT
GCACCTCTCTCCTTCCAAGCCCTCTCCCTTCTCTTCTGCTGCTGCCTGT
CATGTCAAGCATATGCATCAGTGGCAGGACATTTGAAATGCAACCAG
TACAATTGGGCGCGGTTATGCCTACCAGTTTTCTTCTTAAACATTTTA
TATTATGTTTGAAGCATGCCACCTTTCTTCACTTGCCAACCTTGACAGA
TTTATTAGTTTGACAACATCCGCTGATAGCATCAGTAATAAGTTAATTGTT
TTTGCACTGTAGCTTTAATTATTCTCATTATCATTTATAGGAGTTATTC
TTTGTAAGGGTAACTGAGTTTTCAAAAACAAACAGAAATTTGGGGTGGG
CCCATGGAGCGTGACTCATGAAATCAGATTCTTAGAAGGACCTCGGCAAG
TCTCTGGGTTGCTGTTAATGAGCCTGGCTGGCTGCCAGGGGTGTGTCTGC
CCTTTATGAGGCCACCACTGTTCAAATGCTTGCTGCAGCATTACTTGCC
TAGGTAGTGTCTGTTTCTACTGAACTGTGAGGATCCAATTCTTTGTGGT
CTAAGTAAACAATACTCAGATTCACAAGGAATTGATTAATAAGCCAGAATG
CCAAATGATTACATTTTGTGTAAGACCATATTTACAGTGATTGTATCTG
CTCAAGCTCAAATTAGGATTAGAGTTCTGACAAATACATATGTGAGAAGT
ATGAGGTTAAATACTTGAAATTTGGACTTTTCTAGAAAATCTGAATGTGA
TTGCCATTACATACCTTTCTGGGGATGATGATTCTTGACTTTTATTTT
AAAAGACATAGAAAATACTTAAGAATCAGATTGCTTGGCTGGGCACAG
TGGCTCATGCCTGTAATGCCAGCACTTTGGGAGGCCAAGGTGAGTGGATT
GCTTGAGCTCAGGAGTTTGAGATCAGCCTGGGCAACATGGTGAAATCCCA
TCTCTACCAAAAATACAAAAAACAACCAAAAAGAATAAA
TTAGCTAGGTGTGATGGTGCCTGCTTGTAGTTCCAGCTACTTGGGAGGAT
GAGGTGGAAGAATTGCTTGAGCCCAGGAGGTGGAGGTTTCAGTGAGCTGG
GGTTGCAACAGTGTACTCCAGCCTGGGCGATAGAGTGAGACTCCGTCTCA
AAAAAATAAATCAGATTGCTTTATTGCTGGTTTTCTTTCTAAACTGA
GATTGGGTCCCATCATCCCTGGCCCCCATTGGTTAATGGTTCTCCTTT
GTCTATTGAATAAAATACAGATGTCTGCTTTTGGCAACATGGTTGAATGT
AGACACTGCAGGCTCTTCTGACTCAAAATGAGTAAGGCTTAGATAAAAC
ACATTTTGAAATGCATTTCTGGATGAACAGCAAGGAAAGGAGATCTCTTA
AAATCCTCTTTCTGTTCCCTCTCCCTACCCCTCCAAGTGGCTTAAGT
AGGAAGGGTGGTGAGCGGCAGGTAAACACACGTCAAAGGCAGTCTTCCTC
TCTGAGGGAACACTTGTATAAGCATTGCAATCAATGGGCTCTTTAAT
TATGTGCCAGTGGCAAGAGCGGGTGTGAACCCAGGGGCTGCCTCAATC
CGGGGCTTTGAGGCAGAAATAAGTGGTCTCAGGTGTGTGGCATTCTCTT
GCCCTTCCACCCGAAGCAGACACAAATCCTCTCTGGAGGCAAGTTCCCA
ATTGAGCCAGTACAACCTCCACAGACTAAGATCAATCATGTACAAGCTCA
CAGACAAAGGTCAACCAACACACAGAGCAATAAACAATTCATGAGTGAC

GTGAATGAGAATAAACA...AACAATAACCACCAGCTGGGATGCTCTAAG.
CTTCAGCTGTTAGAATTCTGAATATAGAATAAACTGCCACAATGGCAA
ACATGCATCTAGTACTTACTGTGTGCTGGGTTCTAAGAATTTTGCACAT
GTGCCAGATACCGACTCAGCTTCACACTCACCTCTCTACTGTGCCCTCT
AATTTGCAGTAGATTAAAAGGTAGAAAGGAAGAGGCAGCTATTCTGTTCT
TGGCTGTGCCCTCTGGCAGCACATGCAAAATGGGCAGTAACAGTGGCAGTC
ACAGGTAAGTAGCCTTCTCACAGTGTGGAGTTAAAGGCATGGGACTGAGA
CGAGCAAGGTTCTTAAAGGGACAGTGGCCAGTAGATGACCAGGGGCTACT
GGAGTGGCTGCATGGCTCTGTGGAAGCTCAGAGGAGCCTTGGGTCTTGCA
GGTGCAGTAGCAGCTTCTGTAGTTCCTGATCTCTGGGTCCCACAATCTT
CCCCGTTTTTGTCTCTCCACTTCTAATTTTGTAACTGACTTCCCTGTGTG
TACTTCTCTCTGTGATTGAAATAGCCAGACTGGTTTCTGTTTCTTGATAA
GACATTGTCTGGTACGAACACAGTAACCTATTTAATCCGATATCTCTATG
AAGGAGGTACAATAATTATCTATTTTACAGATGAGGAAACACAGCAGA
GAAATAAAGTCAATTGTCTAAGGTTGCACATTTAGTCAAGGGAAGGGTTG
ATATAACATATAATTATTTAGAAAACATCTAAGGAAATAAAAGGCATAAT
TAAAAATAAACTAGGCAGGTTTAAAAAATGAAGTAATCTATAAGTAA
AAAAGTATAATTGTTGAAATACATATCTTAGTGGATGGGTTAAATAGCTG
AAGAAATGATTAAATGAACCTGGAAGGTAGTCTGAGGAAATCAGAATTCAG
CATAGATAGAAAAAATGGGAATTTACAAAAGTACACAGGAATTATAAAG
AGGTTAAATTATAGGGAGGGTAGAATGAGAATTAACATTGGTCTAAGTGG
AATTTTGAAGAAGAGAATAGAGAGAATGAACAAGGCAATATTTAAAGAG
GTGGCTGAGAAATTTTTCAGAACCAACACAACTATGACTTTACCAGTAGA
GAAAACAATGTACACTGAGGAGGATAAATAAATATACTATGAACAAATTG
TAATAATAACTCAACAAAGACAAAGAGAAGATGTTAAATCAGCAAAA
AAAGAAAGTCAGACTTAGAAAAGAAATGCAATGGCAGACTACTCAACAAC
AACAAATGGAATCCAAATTCGGTCAAACAGTATTTCTTCATGCTAGCATA
TAGC

>Contig40

GGGAGTCCGCTATGCTCCTAAAGATTTGCACCTCTGATCTGGTTTGTAGT
TAGTCTCTTTTATGCTTTATCCTACTCAACTAATTTTTTTAGTGCCTGT
TTTTTTTTTTTAAATGTGTGTTGATGACTACAATCTAAACTCATTCTA
CTGATTATGCGGTCTTTAAATCTGAGCAGTCTTTCGCATTTACTGCCT
GTGATGGCCCATCCCACAGCTAAAGTGTGTGGCCACTGCTTACAGCACC
ATGTGATAACGAGTAAGGGAGAGATGCCGCCAGACTCTTCTAGGAGCAG
CCAGTAGGACCTTCCAGGGGTTGCAAGCAAACACAGCAATATGTGGAGT
GTGGCAGAGSATGGCCCAAGAGGATGTGGCAGCGGCTAGTGCAGCTCAG
CTTAGTCTGAGAGGAAATGCTGGAGAGGAGAGCCAGTCTGTACAGGCAT
GACAGCCCAAGGACTTCAACAGCTAACATGGCTGAGTGGACTTTATGTG
CTATCTCATTACAGAAAACAGGAGCAATCAGAAAGGAGTCACCTCCTATTT
GTACCCAGGAATCTTAACCTACTTGCATCTGAATGATGTCCATCACTT
CCCTTCATCACCTTCTCTGGGGGCTCTGCAAGGATTTGACTCCTGCATTA
GTGATCTGTCTCACCTACGTTGTGATTACATGAACCTTACTAATGTGCTA
TGTGACAACTACCATCTTAAACACAAAACCTCTTTTGATTCTGTGGCT
CCCTCCAGCTACCCCTGCATTTCTCTGTCCCCCTGCCCGTCTCTGCACT
CACTTTTATTTTACAGCAAACTACTCAAGGGAGTCTCAGTGTCTCCTTGG
CTCCATGTCTCCACCTTTCACTCTCTCTCAGTTCACCTCTGTGAGGCTT
CCGTCTCAAGCTCTTCTTCACTTTTGTCTTAGGGCCGCTGACATCCTCT
TTCTTGCCAAATTCAGTGGCCAGGTCTCACTTACTCAACTGCTCAGCAT
TGTGAGGCTGGTGGACCACATTCTCCTTCAACCCTTTTGTGCTCTC
TCTTCTCTCCAGATGTTTCTCTCTCTCACTGGCTACTCCTCTTTTGTCT
CCTTTGTAGTCCATTTCTTCTTCCAACCTCACTGTGCTGGTGTGCCC
AGTGGCTCAGTTTATAGCTATTCTCTCTTTTCCAGTGGCATTATTAGATG
GTATCATGTGACCCATGGCATTATATGCCTTCTACATGACAGTTACTCCT
GAATATGAATCTCAGGAAAGATTTGGATTTATTTTAAATTAATTTTTTA
AATTTTAAATTAATAATGAGGTCTCTCTGTGTCATCCAGGCTGGAGTGT
AGTATTGAGTGATGTGATTATAGCTCACTGCAGCCTTGAACCATGGGCTC
AAGTGATCCTCCTGCCTCAGCTTCTGAGTAGCTGGGACTACAGGCATGT
GCCACCATGCCTGGATGACTTTTTTGTGTGTGTGTGTGTGTGTGGAGACAG
GGTCTTCTCTATTGCCAGGCTGATCAAACTCCTGGCCTCAAGTGAT

FIG. 3 (39 of 52)

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CCTCTCACCTCAGCCTTCAAAGTGCTGGGATTACAGGTGTGAGACCACTGGGCTAAGATTGAGATTTTGTATTCAATTGACTGTTTGACATCTTCAC
TTGGACACCTAAGAGGTATCTCAAAATATTAATTAACCTTGGCCAAAATACA
GAACCTTTTGACCCCTGCCCCACAATACTTGCCCTTCCCCAGACTTCTC
CAATTTCTGTTAAATATCCCCAGTTACTCAACCTCAAACCTATGAATGCC
CTTTGATTTCTTTCTTTCCCTCATCTCTACGTTGACGCCATCAGCTAGT
TTTGTTCCTTTATGCCCAGAATATAATCCTCACCACCTTCTCTCTATT
GCCCGAGTATAAGATGTCAGTTTTTCTGACAGTCCATTGCCCTGACCT
CCTGAGTGGTTTGGCTTCCACTTTTGACATTTGTATTCTCTTTCCCCCAG
GGTCAATTTTTCACAGCAAGAGTGGCATTTTTTTTTTTTTTTTTTTTTG
AGACGGAGTCTCGCTCTGTGCCCCAGGCCGACTGCGGACTGCAGTGGCG
CAATCTCGGCTCACTGCAAGCTCCGCCCTCCCGGTTTACGCCATTCTCCT
GCCTCAGCCTCCCGAGTAGCTGGGAATACAGGCGCCCGCCACCGCGCCCG
GCTAAATTTTTTGTATTTTTAGTAGAGACGGGGTTTACCTTGTAGCCAG
GATGGTCTCGATCTCCTGACCTCATGATCCACCCGCTCGGCCTCCCAA
GTGCTGGGATTACAGGCGTGAGCCACCGCGCCCGCCAAGAGTGGCATT
TTAAAACCATATATTAGATCATTGCTTTTGTGTTTGGGAACCTCCAAGGG
CTTTGCATCATATATCAAGTTGACACCTCTCTACCCAAGCCTGGCTCTT
TCCTGCTCCTCTGTCTCTCAGCCCCCTCCACCCATTGTTTCATGCTGCTTC
AGCCACCATGGCCTTCTTGCCATGCCACATTTGTGCTAAGCCACATCCA
ATCTCGGGGCTTTGCACTCGCATTTCTCTGCTTGGCATGCTGTACCCC
AGATCTTTTCATGATTGGCAGCTTCTGTACATTCAGCCACCTGCTCAAGCC
ACCCCTTTCAGAGGGCCTTCCCTGGCCACCTCACCTGAAATAGCACCTCCG
ATTGCACCCATCCGGTTATTCTCCATCCTGTTCTCTTGGCTGGTGATTTT
CCATCACTGATGAGGAAATGAACCATGGAATGCTAGGGCTGATGACCAGA
ACTTTCCCCCACCCTTACATTATACAGAGGAGGAAATGAGGTCCGAGGT
AAGATGGGCCCAGGATTTCTACTCCCGCTGGACTGCAGGCACAGCACTG
ACCTCAGCTGTGTCACTCTTGCAATTCACCCAACCTTCTATCTCCAAC
TGCCCCATTTACCAAGAGTGAATGTTCTCAGAGACGGTGAGCCACCTG
ACTTGGACAGCAGCCCAGGGCCCCCTGGCACCTGCTTTCTTCTCCTGC
CATCCTTTCTCTCCAAGACCTACCTTTCCCTGTGATTCTTGCCACATG
CTGCATTTTCATGGTTTTATGACCTGATTTCTGAGAGGGATTTGAATTTTC
ATGATTATTTATGTAAGCAAATCATTATGCTTATACAAATGAGAAAAGGA
GTGCTTCTGGACTTCCAGGGACAAAATCTTGTCACTTGGCTTGGCTTCA
TATTGCTAATTAAGGACCAGGATGTGGGTGAGATGTGCTAAAAGCTGAG
AGGAGGCTTGGACTCTGACTATGGGCCACACCCCTGGGCAGGCATCAC
ACTAGTCTTTAGGTCTCTCAACCCAGCTTCCAGTTGAATCAGATGTT
TGTGAATAACTCAGCAAGGCTGTATGGGAAATGAAGAATGAGGTGGGGAA
GAGGCTTGTGAGAGACACACTGACTTACCCCTCTACCTCTAACTAGGG
TGTGTAGCAGCCACCCACCCACCAAGTCTGTCTTCCAGACCAGTATGC
TTTCTCCACCTTTGCATCTTTTATCTTCTGCCAGCCAGATGCTTGCTG
ACTCCAGCCCAAGCCTATAGGATAAGCTACAGCCTGTCCCTACAGACTAC
GCATTGCAGAAATCTAAGACATCAAGTCAAGTTCCGAAGCACTTGCCTTCT
CCTCTCCAGGTACACAGGCTCTCCTGGAAAGCTGGTAGCAGCTGTGGAGG
TGTGGTGTGTACCTGCTGCAGGTGCAGAGAAGTTGACTTCACAGCCCTT
CAGAAAGACTGCCTTCTTCCAGTTGTATTGTGTACTTGTGCTGGGTGTGG
GGAGGATTCTCAGCTTTCTCACTCAAATTATCAGACCCCTTCCATTTAG
TGGTAGACCATTTCCCTCGTCCAGGCCAAGGGCACATAGTACAGAGAAAT
AGGGAGTTGTTACCCAGGGAGAGAACTTGGCTCTAAACCTGTAATAGAAA
GGTCAGTTCTGGTCTGGAGGGTCAATTTTGATCTTTGGCTCAGATCCAGG
AATTGGAACCAAGGCTTTTGAACATTTTAATGCAGGGGATTAAAAAATG
ATACGAGTCATTACGAATATATTTGCTTAACATCTAAAGAGATCCCTCA
AAACACTAGAAAAAATAAGAACAAAAATCTAATAAAACAAAAATTTGTAA
ACACATTTACCAAAATTTTTTTTTTTGGTAAAAATCAAATGTCATAAATA
AAGCTAAAGTTCTCTTGATGACTCGCTCCTCTGCCCTATTCCACTCCAA
GTAACCACTATTATCAGTCTTGCCAATACCCTTCCAGACCTCTCTACCTC
TATATACCATTAGAAGCACATGGTTTTGTCATTGAGGATGTGCAGTGTGTT
GTTTTACGTAAATGTTATCACTCTGTTCTTGTTCATAATTTGCCTTTTT
CTCTCAATGATTTGCTTGGCTATCTTTCTATTTTCACTAGCATCTCCTTTC
TTTTTAACCTTACCATTTGTTTATTTAACCTTGCCTCTATCAACAGATATGT

FIG. 3 (40 of 52)

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AGGTTGTTTCTAGTTGA.TTCATTAAGTATTTATAAACAACGCATCAGTA
 BATGTCCATAAAATTTCTTTACGGAAGATGGCAAGTAGTGGAAATTTGCTGAG
 CCAAAGAACATGTTTTAAAAAACCCTAAAAAACTAGACGCTACCAATTTTC
 TCTCCAAAATGGCCATACCCACTTACCCATACAGAGATGATTTGGAATCT
 GGCTTCCTCACAAGGTGAGATGCCTTCACAGTTTCATTCTTCTGGCATG
 TCTTCCCTTTTGTATCTGAGAGAGCTGGCAGAATTGTGTCACTAAATCAA
 GGATAGAGGGTCAAATGACAGCTCAAGCTCACAGGCACCTCTGCTTTCTT
 CCCAGACCACCTGCTTTCTGCCCAGCTCTGTTCCATCTTATAGAATG
 GTTGCCACTTGGGTGCTGCTCCGACAGCCATGTCTCTTTGCACTGCA
 GTTATGAAGCAGACAGAGCTAGGAGAGGGGCTTTGCCAGCCTCTGCCCTA
 GCTTGGAGAATTTCAAAGAAGGAGGGTATTGAGAGTGAGCTGCCGAAGAC
 TGGCAGCTCCCTCAACTCAACAGTTGTCTTCCACAAGAAGTCAGATACA
 TTTTTTTGGGATAAAATATTTAAAAATTATTATTTTATTTCTGAATAATA
 TATTTACATGATTTCAAATACTAACTGTAGGCCAGGCATGGCTGCTTATG
 CCTGTAATCCTAGCAATTTAGGAGGCCGAGGCGGGAGGATCACTTCAGCC
 CAGGAGTTCAAGACCAGCCTGGGTAACATAGTGAGACCCTGTATCTACAA
 AAATTTAAAAACAATAATAGTTGGGCATGGTGGCTGATATGGTTTGGCT
 CTGTGACCCAACCTCAAACTCATGTTGAATTTAATCCTCAATGTTGAGG
 GAGGGTCTGGTGGGAGGTGATTGGATCATGGGGTGGGTTCTCCCTTGC
 TGTCTCATGATAGTGAGTGAGTTCTCACAAGACCTGGTTATTTGAAAGT
 GTGTAGCACCTCCCCCTCACTCTCTCACTCTCTGCTCCGCCATAGTAA
 GATGTGTGTGTTTCCCTTTGCTTCCGCCATGATTGTAAGTTTCTTGAA
 CCTCCAGCTATGCTTCTGTACAGCCTGTAGAATCTGTAATCAGTTAG
 ACCTCTTTCTTCAATAATACCCAGTCTCAGGTCATTCTTTATAGCAGT
 GTGAGAGTGGATGAATATAGTGCCATATGTTTGTATTTCCAGCTACCCAG
 GAGGCTGAGGTAAGAGGATTGCTTGAGCCTGGGAGTTTAAGGCTGCAGTG
 AGCCATGACTGTACCACTGCTCTCCAGCCTGGGTGACAGCGAGACCTTGT
 CTCCAAAAAATAAACCCTAACTGTGTAATAATGTGTTCAATAAAGTGTC
 TTGCTCCACACCTGTCCCTATATATCTTATCTCAGCCTCCGACAACCT
 ACTTTATTCAATTTCTTATGTATCTTCCAGAATCAAAAAAATAAATAA
 TACAAGCACAGTGAATGTATTGCCCTTCTTCCCTCCCTTTTGTACAT
 CAGAGTTAGCATGAATATACGGTCTGCATTTCTTCTTTTTCAGCTA
 TCAGCATGTTTTGGAGAGGATTTCAATTCGTGCAGACAGCATGTATTAG
 TCAGTCTTTCATTGCTATAAGGAAATACCTGAGACTGCATAATTTATAA
 AGAAAAGAGGTTTAATTGGCTCACAGCTTCGCAGGCTGTTCCACAGGAAG
 CATGGCAGCATCTGCTTCTGGGGAGGCCTTAGGAAGCTTTTACTCATGCA
 GAAGACAAAGCGGGAGTGATGTTTATATGGCAGGAGCAGGACTGAGAG
 AGAGAGAGAGAGAGAAAGGATGCCACATACTTTTAAACAACCAGATCT
 TGTGGGAACCTCTGTCACGAGAACAGCACCAAGGGATAGTGCTAAACCAT
 TCATAAGAACTCCACCCCATGATCCAATCACCCACACCAGGCCCCACC
 TCCAACATCGGGGATTACAATTTGACATGAGATTTGGGCTGGGACACAGA
 ACCAAACAATACCAGAGTGCTTTCTCATTCTTTTCTATAGCTGCCTAGTA
 TTCTATGTCTTTTACTTCAATTTAGGCAGTCTCTTGTGATAGACACTTGG
 GTTACTTCCAATTTTCTTATTACAAATGATGTGCAATGAATAATTTTGA
 TCATTTTCCATTTCAATGGGTTATGTCCATCTGTGGGATAAATCTCCAG
 GAGTGAATTTGCTGGATCAAAGGGGAAGTGCACTTGTGATTTTCATAGTT
 AGCAATTTTGTCTATAAGGGTCATATCAATTTATAGTCCCACGCGTAA
 TATTTAACAGTGGGGATTTCCCGACAGTTTGACCAACAAGGTCTGTTGTT
 AAATTTTGTATTTTGTCAATCTGATGGGAAAATACTAGTATCTCAAAGT
 GCTTTTAAATTTGACTTTCTTATTACAATGTTAAGCATCATTTTACTCTGC
 CCAAGATCAAATAGTATTTTCTTTTCTGTGAACAGACTGTTAAGATCCCT
 TGCCTCTTGTGTTTGTGCTGGATTTTGTCTTTTCTTTTCAAATGTTTGTAGG
 CAGTTCTTTACATGTGAAACAAGTTATCTCTTTATCTGGGGTGTGAGTTA
 CAACTACTTTTCTCTGGCTTGTGTTGCGCTTTGACTTTGCTTCTGGTGA
 TTCCCGCAATTTCTGAAAGTGTAATTTTGCATCATTCAATCTTATACACC
 CATGCTCTTGTTCACGCTGGTTCCTCTACCTGAGGGCTTTTCTTTTCTG
 CTTCTATCTGGGAACATTTTTTTGAGAGAGAGTCTCACTCTCTCGCCAG
 GCTGGAGTAGTGCAATGGCGCGATCTTAGCTCACTGCAACCTCCACCTCC
 TGGGTTCAAGCAATTTCTCTGCTCAGCCTCCCAAGTAGCTGGGATTACA
 GGAGCCCAACCAAGCCAGCTAATTTGTTGATTTATTTATTTATTTT

FIG. 3 (41 of 52)

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TGTAGAGATGGGAGTC. LACTATGTTGCCAGGCTGGTCTTGAACCTC. J
GGCTCAAGCGATCCACCCACCTCGGCCACCCAAAGTGCTGGGATTACAGG
CCTAAGCCACCATGCCCCAGCCCATGTGTGGAAATCTTCTGTTTATCCCTT
TAGGCTTGATTCTTATGTCTGTTCTCCTCCCTCCTTCTGGATACTCCTCT
TGTCTCTTACTCTTACTCTACTTGTCTGTTACCTGTTTCTGCTTATAAC
TAGCTGCTCTCTCTATCTGAGGAGGACTTGTGACTGTTCTCATCTCTGT
ACTCCAGCTCCTAGTACATAGCGCTTGCTCAACAGATGTTTGGTGCATT
GATAGATAAATCACTGGTAGCTGTTACTACCAGTCCTGACTCCCTGCAGT
GCTTCAGCTGATCCTGTTCCAGATGTGCACTGAATATCCTTCTGTTGAAC
AACAGAAATAAAGGGGATGGGTGAGGAGGATAGTCTTCGGTGGCCAAGGA
TATTTTTAGGTACTTTGCAGCACTCAGCAATGAGGAGTGGGCTTTAGTCC
CCCAAGAACTCTCACAGCCCTGGGTGCTTTACTGTTCACTGTCAAATCC
AAGACAAGTCAATGATCAGGAAAGACCAATTTTTTTTGTTCAGTGAAGTT
TATTTTCAAGATCATTGAACAGTATGATATTTGGTAATTTCAATAATATC
CCACTTAAATGATCGGAGCAGATATATTTTCACTCGTAATTAAGGACA
TGATTTAAAGAGAGCACACCAGTCCAAATTGAAATGATTCCATAGCTATT
AAAAAACTAGGGTTTTTACAGACAATGATACTTTTTGCCCTTTGAAT
AGATTAGACCAATGAATAAAACAAACAAATAAAATAAATAAATAGGG
AAGCGGTTGCTCATCAGAATGTGGGAGCGAATGACAGAGGGTTTCTTAGA
ACCAATATGTGGCCGTGTTCTGTGTCAGGCGTGCTTAAAGTAGTAGGAGA
GGTGAGAGGGCCTGGCTCAACAAAAGGGCTGGGGATTGTCCCTGAAGAA
CCAGAGCTGANTTNCATCAGGAGTAACANAGGTAGATAG

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CCGCGTTGAGGTTCCACGCAGTTCAAATTATGTCCAATTATCAACATTAA
TGCACATTTTCAATAGAACCTGTTCCGGCTTTTCTTAGGAGGGGGCGGG
GAGACGTTGTTCTCTGGGAATAAGTGTACGCAGGAGGCTGAGAAGGCTTC
ATTCCATAGCATTCACTTACCTCCAGCTGTAGAGTGGGCTTATCATCTTT
CAACAGCGCAGGACAGGTACAGATTTTTTTCTTTGAGGCCAAGGCCACAG
GTATTTTGTCTTACTTTCTTCTCTTGTACAAAGGACATGGAGAACC
ACTGAAGAAAGAAGGGGGTCTTGTGGTTAGGGACACAGCAGTGCAGGGTC
ACCCCAACCCCTAGGCCCATGAGTAGGATACATGTAATTTGGTAGCCTC
TGTGGGAACCCACAGTGAGGTTCTTGGCCTAAGACACAGGATAACTTGA
CTTCTCACAGACAATAGCAGGGTCATTTTGTGATTAGGGTTTCCCTC
AAAGGCTGAGGGTTTCTCAGAGCCTCATAGCAGTAGGAACGGAGAATGA
AAGAGGCTCTACATTTTAAATGCTGAAGGAAGGAAGGAAGGCCATTG
TGTCAGTGGCTGGCAATGTGCCCATCCACAGGAGCGGAACAACTTGATCA
ATGTGGAAGGAAGGAAGAGGTGAGGCTGTACTTCTGCCAGAAATCAGG
CACCAGAAGCTTTTTAGGAACAGAGAGTAGCCCATGGGAAGAACTGGGA
GAGGAGAGGCTGAGCTGGGAAAGTGGCTCCAAAGAGAGACACTCATTTG
ATCTTCTCAGTCACAGCAGTGTCAATTGGAGGCCCTGGGATCACTCTTA
CTACCCGATTCCAAAGAAACAGGATTTTCTTGGCCTGGCTGAGAGCAAAT
AGCTTCCCTGAGTGAGGCTGTCTTCAAAGTCAGCAGCCTTAGTTGCC
CACACTCCTGTGCAGAGGCTTTGGCTACTGTGGCACGATGCCAGGCAGAT
CACCACAGCTAATGATGGGTTCAACGCACTTGAACTTTTGCCCGTTACA
GCGGAGAGATATAAGTTCTGTGCTGGGCGTAAATTTCCCTACAAGGAAC
CACCTGGCATTGGGTGGGACGGATGTTGGGGCAAGGGGGGAAGACTGGGG
AGGGGGATGGACACATTATCGCTCCAGCACTCTTGTTCAGCCTCAACAA
CAGGAAGAGAGAACCCACAGGCAGTTAGGCCATGTCCATCAAATGACCCC
ATATTGTGGAAGAATTGACATTGCACTATGCCCAAGAGACTTGGGTGGAC
ATGGTCTCTGGGAGTGCTTGAGCCGTCTAATTTCTCAGGGTCACTCCTG
TTAACAAATGCACTGGCCAGTGCAATCAAATGTGCCATTTCTAGGACCAA
AGTTTGTATATTCCTTTTTAATATTTTTTTCACTTGTGTTGATCATTG
CCTTAAATTAACCTTTCTACTTTGTTTAAACATGGAGAATTAGCAAGCTG
CCAGGAGGCCAGGCAGGGAACAGGATGTTTCCATTTACCTTGTGCTC
CATATCCTGTCCCTGGAGGTGGAGAGCTTTCAGTTCATATGGACCAGACA
TCACCAAGCTTTTTGCTGTGAGTCCCGAGCGTGCAGTTTCACTGATCGT
ACAGGTGCATCGTGCACATAAGCTTCGTTATCCCATGTGTGGAAGAAGAT
AGGTCTGAAATGTGGAGCACATGTTGTTTAGGTATAAAATCAGAAGGGC
AGGCCTCTGAGGCGAGGTGGCAAAATTTGATTTCTTGGAGGACACCTGA
GCATATACGGTCAAAGTCTGATGACAACACCAGTAGGGATGAAGCTGGGA

FIG. 3 (42 of 52)

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GTGGGGTGGCTAAGAACTGGACCTGACACTATTAGACATGGGTTCC...
CTTCAGGTCTATTACTGCTCACTGTGGCCGAGCAACAGAGCTACTTAGGT
AAAATGGTGTATGGTCATAACACTAGCCACAGGGAGGTTACGAACCTCTG
GTGACAATGTAAAGTGAAAGGCCCTGAGAAAGAGTGAGGGGAGTTGCAAAAT
GTCAGTAGCCATCAAGATCTTCTTTAAGAATAGTTTCCACTAAAGAGATG
ATTGCTTTGGTTTCCAGCCTTCTTTGTTTTGTCTCCCGCTGGGCCTTCT
ACCTTTAAAGGGCTTTGGCTCTGGGGGAATTGAGTTGGCTGGGGCTTGAT
GACTTCCAAGAGGACACAAGTGGAGATCTACTGCCTGCTCTTGGCTAACT
ACCTTCTTCAAAGATGAAGGGAAAGAAGGTGCTCAGGTCAATTCTCCTGGA
AGGTCTGTGGGCAGGGAACCAGCATCTTCTCAGCTTGTCCATGGCCACA
ACAACCTGACGCGGCTGCTGAAGCCCTTGCTGTAGTGGTGGTGGGAGAT
TCGTAGCTGATGCCGCCATCCAGAGGGCAGAGGTCCAGGTCTTGGAAAGG
AGCACTGCGGAGAGAGCGAGGGAGGGAGCCTGGTGAGGTGGTCTGCCAG
GAACCATGCTTTGACATCAGAGAGTAGAAAGCTCAGAGAGGAGGAAAGGG
CTTGAAAGAATCCCGAGCTTCTAAAGATCATCCCTCTCTGGGCCAGGCGT
GGTGGCTCATGCCTGTAATCCAGCACTTTGGGAAGCCGAGGTGGATGAA
TCATTTAGGTGAGGACTTCAAAACCAGCCTGGCCAACATGGCGAAACCCC
TTCTCTACTAAAAATACAAAAATTAGCTGGGTGTGGTGGGGTGCACCTGT
AATCCTAGCTATTGAGGAGCTGAGGAAGGAGAATCGCTTGAACCTCAGGA
GGTGGAGGATGCAGTAAGCCAAGATTGTACCACTGCCTCCAGCCTGGGC
AACAGAGTGAGACTCTGTCTCATAAAACAAAAACAAAAACAAAAACAA
AATAAAATAAAATAAAATAAAAGATTATCCCTCTCTGAAGCTCAAGGAG
GTTAAGGGTGTACTCAAGGGGCACACAGCAGGTTAGAGGCAGACTCAAGAT
TAGAATGTGGGCTTTCTGACACCTTACAGGCTATTCTTTTAGAATAAATC
CCATTTCTACTTTGTTTCTCTTTTGTACATGCCCCACCTACACCATAC
ATGTATACCTTCTCTATATCTTTTGTATCCCTAATGCTGTCACTATG
ATTTGCTTTTCTATGCAGATGACCATAACATTTCCATTACCTATGCTC
ACTCAGCAAGTATTCAATTTTCTACACTGTTCTTTTTTCTTTTTTCA
TAACACTGTCTCATAGGCATTCTGCAAATCCTGTGAGAGTACTTTTTGTG
AAATGTTACCACTTTCTCTTATTGAGAGAAGCTCCGTATTAAGGCTTCA
CTGAGGTTGCCTTAAGGCATGATAATGGTTCAAAGGCTTGAAAGACAGTT
AAAGAGACCTGTAAGTGACAAAAAGAAAGTTGAGCAGGAGAGAATTTCT
GCCTGGAGCAGAGCCAAGCTGCTGGAAGAGGCAATGGGGGCAAAGGCCAG
GCAGACAAGCCAATGGGCTCCTCCACAGCTGCAGCCAACAAGTTATGCC
AGTCTTAAACCTTCTAAAGAAATATGTTTTTAACAAGATTGAGGACTGGA
TTATGAGGCTAGGGGAGGCTATCACAACCTGGAATAAAATAAAGCCAGAG
AAAAGTGGCTGCCTTCCAACCTGCACAACCTGACCTAGCTAGGCTGATGGC
TGGGCCACCTAGGAAGGCTACTGAGCATCATATAAAACAGAAGGGACAGC
AGGAATATAACATGGCTCTTTGTAAGGATGAGTCTGAAAAATGACCATT
GCTGCCCAAATGCCCTTAGCTACAACCTGAAATATTTTCAAACTGGAGGT
TGCAGGATGCTGGAATCTCAGAGATCATCCAGCTCAGCCCTTTATTTTTT
AGATGAGGTCCAAAGCGGGTAAATGACTTGTCAAGGTCAAACAGCAAGT
GAATGGTTTTCTTCAAGTCTCAATTCATTTTTTGTATATCATCTAT
GTCTTGTGTTATAAGCTTCAACCCAGGTAGCAAAAACTATTCTACTCA
AAAGGGGTAGACATATGTTAGTTCTCAAGATCATCTTGGTTTCAGAGT
TTAACTCAAGTGATTGGCATAGGCTGAATCCATCTCTTAAAGGATAATC
AAATTTATGTTGAAGACTTGGTTGTCTTCTACTATGAAATGGGAAACAT
TATCACTACTCCTCCCTGTCAACCAAGTGTGGCCACCACCACCAACG
TTAGTGAGTGACTGTGGTGATATGATGACCAAGTGGCCAGGTCAGCAAGT
GGTGCAGCCTGTGTCTCACTGGAAGAGGTTAAAGTCTTTCTAAACAAAA
TACCATGGCATCAAAGTGGCCAGAACTCCCTTCTTTGAGCTTTCCCTGT
GTTAGAGCCCTTCCCTGGGTTGGGAGTTAAACCCATAGTCTTACCTTCAT
CTGTTTAAAGGCCATCAGCTTCAAAGAACAAGTCATCCTCATTGCCACTGT
AATAAAAAACAGGACATGTCTCAATTATGTCTTCTAAACAGGTTTATTTT
TCCTTCCCTGTGTACAAGACTTGACTGTTTATAAGAACTGCAACAGCC
TGCCTCTCAAAGCTGCCTGAAACACCTGGCAAGTTTACAGTGATATGCG
CAGAACAGTCCAGAAGGCAGATTCTAGGCCTGGCAGGTGGGCACCCCTGGG
TGCTCCCTGTGGATCTTGAGGCCTAACCTCTAGCCAGCAGAGTCAGCT
AAAACTGAGCTCTCCCTCTCCCTCCAAGCCACACTTTGCAAAGGGATTC
CTTGATTTGGGCTTGAATCTTTTTCTCCCAATTTGCCTCTGCAGGAAG

FIG. 3 (43 of 52)

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CCCTTSCAACACACA TGGATAGCCTCCAGGTCCCAAGGCTGGAGC A
CTTGTAATGGGAAAGTAGTCTTTAAATCAGATTTACTTGGCACCCCTGTTT
GCCACTGAAAGAGGCAATTTAGGGGAAAAATCTGGTCTCCAAGCACAGAT
AACACTCTACTCTTGAAAGAGGAGACCTGCTCATGTTACTGGTCTCAGCG
TCTCCACTGACCTGTAATAAGCCATCATTTCACTGGCGAGCTCAGGTACT
TCTGCCATGGCTGCTTCAGACACCTGTGTAAAAAGGAGAAAAAGAGTGAC
TTCCCATGACGGCTACGTTTCATGTGTGATTTCTCTCAGCATCCAGTGCA
TGGCAGTCATGCAAAGAAATGATCTCTGAGTAAATGAATGAATGTGTGAA
AGAGAAGTCCTTTGGGTCTAGAGAAAAGCATTGCTAAACCAAACCCCAA
CTAGCAATGTATTGGCTAGGAGAGCTGGAGCAGAGGCTTTGACACTAACC
TTTAGGGTGTCTAGCTGTTAGATAAGCAGTATCCATTCCCAGAATATTTCC
CGAGTCATAAGCATTATATTACACCTGGCATTTTTGCAAAAAGCTGAGAG
AGGGAGGACAGAGAGGGAAGGAGAGGAGACAGAGAAAAGAAAGAGAGAG
AGAGAGAGAAATGTCATACACAAAAGAGGCGAGAGAGACAGAGAGACTCC
CTTAGCACCTAGTTGTAAAGGAAGATTAAAGTCATACTTGAGCAATGAAGA
TTGGCTGAAGAGAATCCCAGAGCAGCCTGTTGTGCCTTGTGCCTCGAAGA
GGTTTGGTATCTGCCAGTTTCTCCCTCGCTGTTTTTATAGCTTTCAAAG
CAGAAGTAGGAGGCTGAGAAATTTCTCTGTTGAATACCTGATTTACAAT
CAAGTTAAAGGAAAGGGGAAAAGAGTATTGGTGAAGCTTCTTAGGGGAG
GGGACTAATAAAGTGAATAATTTCTCTGGTTCATGGAAGGGCAAGGAGTA
GCAAACTATGACACATTTTGCAAATGTATCACCATGCAAATATGCATTGT
TTTCTCTGACAAATCGTTGTGAGTTGATGTCCACATTAAAATACTGGATTT
TCCCACGTTAGAAGAATGTTTAAATTTAGTATATGTGGGACAAAGTGGAA
GACACACAGATTTATACATGCACATACTTTTCTTCATTCACTTCTTTGTA
CTTAAGTTTAGGAATCTTCCCACTTACAGATGGATAAATGGGTACAATGA
AGGGCCAATAGCCCTCCCTGTCTGTATTGAGGGTGTGGGTCTCTACCTTG
GGTGTCTTCTCTGCCTCGGGAGCTCTCTGTCAATTGCAGGAGCCTCTGA
GGAGAAAATTGACCTTTCTTGGCTGGGGCAGAGAACATACGGTATGCAGG
GTTCAGGCTCCTGACGGAGTTGGGGCAACCTGGAGATAAGCTCACACAA
CCCTGCAAGACCAGGTGCTGTTACCCTAGCCAATCTCATGGATGAACCAG
ATCAATGCCAGATGAGCTCTGCCTAAAATGATTTTTTGGTGAAGCTCTGAA
AAGTGAATATTTGTTTCTGTAAAGAAATATCCATCTGAGACTCTATCTCTTG
GTAATACCAAGAGTTATCAGTTTCTCTTTAACCGAGACACCAGCAAAGTG
CCTGCTCCAGGGTAATGCCAGGGGAGCCCTCCATTGTAGAATGAATGA
GAGTCCAGGTTATGAACAGTGCCTGGAGTGTAGGAACACCCTCCTTTGCC
TCTTTGACAGGTCTGCATCATAACACTTTTTTTTTTTTTTGGAGACAGAG
TCTCACTCTGTGCGCCAGGCTGGAGTGCAGTGGCAGCATCTCGGCCCCCT
GCAAGTTCGCGCTCCCGGGTTCACACCATTCTCCTGCCTCAGCCTCCCCA
GCAGCTGGGACTACAGGCACCTGCCGCCACGCCCCGCTAATTTTTGTAT
TTTTAGTAGAGACAGGTTTACCATGTTAGCCAGGATGGTCTCGATCTC
CTGACCTTGTGATCTGCCCCCTCGGCCTCCCAAAGTGTGGGATTACAG
GCGTGAGCCACCGTGTCCAGCCTGTAACACTTCTTATAGCACTGAGTTGA
AACCTTGCTCCTCCTGGTTCTCCAGGAACTGAAATCTTTTTGAGCCAA
GTCTAGCACAGTGCCTGGCATGTACATTCAAGTGGTAGAGTTGCTGCTT
GAATGGGTGAATGGGAATTTGACAGCATTTTTATTCAAATTAGTATGTGC
CAGGTATCGTGCTCGCTCTGCATTATCCAAGGGAGTGAGCCTCTGTGCAA
GTATTTGAGACACGAGGGAATAGGTTCTACTGTGGGAAAAAGAGCATT
CATGGACTTGCTCTCAAGCAGCCTTCTGATTTTTAATTTGGCTCCCAGT
ATCTTGATATCAGGAGTCAGTCACAAGAACTCCATCTTTAGTAAGTTATA
TTTTCCACAGGAAATCTAAAAGCTGTTCAACATGTTAGTTTCTGTGAAT
TTGATAAGCCATAATCCATTCTAACACTGAGCCCTCCTGAAATTTGGTG
TCTGGTCTCTGAGATAGCTAAAAGCCCTGTCTGGGTGGCCTAGGGACTCC
TCTGTTTTGCCTCCACAGGATCCACTTTGCAAATTAACCACTGGTTCTCC
CGTTGTAGGAACTGCCACCTTCTCAGAGCCTGTCTTTCTTCTTCTTCTT
CTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTT
CTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTT
TCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCT
CTCCCTCCCTCCCTCTCTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCT
CTTCTCTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTT
CTCCCTCTCTCTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTT

FIG. 3 (44 of 52)

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TCTACCTTTATCCCCCAGCTGGAGTGCAGTGGTACAATCATGCATTCA
TGCATGATCACAGCAGCCTCAAACCCCTCCTCAGAGTCTTTATGCGGCAA
CCAGCAGGGTCTGGAGGGTGGTGGCTCTGTGAACCTCTCCTGACAGAACA
CAGAGATGTCTTTGGTCTGTTGATGTGATTACAAGCTGAACGAAGGAAGA
TCAAAGCCAGTGACAGGAAGGAGATATGCAAGGGACCCGAGCATCAGCT
CTGAGTTAGTCCATTCTGCTTCTGGGACTTGGGATACAGGTCAGAAACCT
TGAGCTTCTACTTCTCCATCTTCCAATTGTAGCATCCAGGACCTCAGAAT
CTGCCAGCTAAGAGGAGCCCTAATGATTGTCTGGTGGGATATGGTGGGAC
CACAGAGATGAAGACATGAATAGCTATTTGAATGTGAACAGCAGACGAAG
AAATCAAGGCTAGGAGGGTGGAAAGTGAATCCTCAATAGCACAGTGTGGT
TGAAGCAGCACTAGTATCCAGGTTGCATGAGCCCCGATGCTTTCGCTCG
AGGGAAATTTTGGAGCCATGGGGCAATGCCCCCTGACGTAACAGTCTCCA
CAGTTCTGCCATGTCTCATCTGGCCCTGTAACTGGACCCAAATCTGCT
ACCATCCCATCCATCTCAGGAAGTGAACCTCTTATGTCAAATAGGTTGT
GCAACGTATGTATCAGATCCTGTCTTCCCAAGGAGACCGCTCAGGCCACA
GCACTTCTTCCGATCCCCAATGAGCAGAAAATATCTCGCTATAAACATA
GTTGGCACTAAGGGAGGGAGTGGAAAGTGAATGATGTAGATGGTGTAT
GTAGCCCCAAGGAAGTGGAAACAGCAGAGATGGGGAGCTGGAAATGCCAG
GATGCTCCAGCTTTTGGGGAATTATTAGCTCTTGAGTCACTAAAGCCTT
TCTCAGCTGCAAGTTCCCTCTTTACCCTGTGAGTCACTTCTTCCAGACAG
GAGACTGACATTTATTCAAAGCAGCAAGTGGCCGTGATACCATCTTGTGTC
TAATCATGGGCTTTCGAGCCAGTTATCAAGGTTGATCTCATCTCATTGGT
CTTCAATCATTTTGAACAAGAAGACAAGCAAAATAATCATGGGTTAGTTC
TTATATTATTGTGTGATGATGATGCTGTTCTTTGTAGTGAGCTG
TTCCTTCTTGTTCACCCCTCTTGCTTAGAACAGAACTAAGCAATCTGCCC
CCAACATTTTCCCCAATTTCCCATCTCATTCTTGGCACTGGCTTCCCTAAT
ATTTGTTCTTATGAGTCAATTTCTTGATCATTTCCATGAGTCCCTCTGG
GATCTTAAAGTATGAAAAATGTTGTGTGTACCCACACCTGTCTTTGTGGA
TATTTCTCTCCTTCCCTTCTGCTTCTGGGATTATTTGGGAATGGGCACT
ATGATTTTATCATATCGCTTCCACTTCTTTATGGCATCATCTCCAATG
GGCTTCTTCTCCCTCTTGGATCCAGGTTCTCAGATTGGGGACATGCAGAG
TCCAAGGAACATTCCATTCTCCTCCCTGGTCTAGAACAAGGAGGGCTTAG
ATATATGAGCAGGTGGCTGGGGCTGGCGAGCTATGTAGTCTCCAATGGCT
TTTCCCTGATGTCGGAGTTGTTATGTGAGTCTGGGAGACCAATAAGACC
TTGTCTTCTTTGGATCCATCAGAAAAAGCCCCCTGGGTGGGTAAAGATGG
ATGGCAGGGCTCTCCTACTCTATGTCTTTTCTCACACCTAGTGGGTATAA
GAGAGGGGACCACAAACAGAGGGGGCTCTGGTACCACTTATCCAGGGTCT
GGAAACATTTTCTGTAAAGGGCCAGATAATAAATGTTTCAGGTACAACCTA
CTCAACCTTGCATCATTTTCAAGAAAGCAGTCAGATAATACATAAATGAAT
GGGTGTGGCTGGACTTGTCTTGGGTCCCTGTCTTATATCATTGTATTA
TATCATTTTTTCTTACATACAAATTTAGAAGCAATACTTAAAAAAAAAAAA
GCCGTCCTTTATTGAGCACCTACTAAGTGCCAGGTACCTTTTTTCCCTC
ATTATCTTATTAATCTTTCATAATAACCTTTAAAGTAGATAATATTGAAC
CATTTGACCTATGCAGAACTGAGGTTGAGACAATAAATTATTTAAGACC
GCACAAACAGTAAATGCTGGAACTACGACTCAAATATGGGTAACTGAAC
CAAAACCAGATCTTTATTTCTCACTTTAATTGTTACATATGTTTATTGC
CTCATCTCCTGTCCACATGGTGGCCATCGGCAGACTCCTTTCTCATTCTC
AGTGATTGAGTGACATTTCTAACTACATTGGCCTGGCAGATTACCTCTG
TCCCCATAATGTTTCCACATTGTCTTTTAGGATTGAGATCCTCTCTGTT
CCCTTGTCTTCCCTCCTTTCTTCTTGGCGGTGACGTGCTGTGTGAATT
TGTTTCTTTCTCCTCTCAGGGTAGTACTGGGACTTTCCAAATCAGGGTTT
TTAGTGATCTCTTCTCCTTTCTGAGTTTCTCCTTATTCCCATTCACT
TTCTCATCTATAAGTGGCAGCTTTGTTGCTGGAGGATTTCTTTGTCTT
TTATTCTTCTTAAAGACTTTGTGCATAACTGTCAAAAGCAATCCCTTGAAG
GTATCTGTCTTGGAAATTGTGTGCTTATGATGCTGAAAAATACTCTCTTC
CTAAAGCTATTATAAATGCT

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GGCTAGCTGCAACTCTTGAATACAAACACATTTCAGACATGCACACACTTT
CTGGCTCCCCAAAAGAAAAAATCAATTTATAATAATTCTGATCCT
TTGCTTATTTCCACAACTCCATGAAAATTGTACATTGTCCAAGCAACAT

FIG. 3 (45 of 52)

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TTCTTAATATCTCTTTA . TCTCTCATATCCATTTTCCTTACTGCTGTC . J
CACCTATCTCTTCCAAACTCCCTGTTAAATCCCTGCCCCAGCGAACTTT
TATTCATTTTGTGGAATGGAGGCTGCACTGATTTAAATTAATAAAAAA
AAAAATCCCTACTCCATGTCCAGATCCCTAGTTGTTTTTTGTTTTTTG
TTTTCTGAGACAGGGTCTTGTGTCTTCCATGCTGGAGTGCAGTGGCATG
ATCATGGCTCACTGCAGCCTCAACCTCCTGGGCTCAAGTAATTCTCTTGC
CTCAGCCTCCCCAGTAGCTGGGAGTTCAAGGTATGTGCTACCATGCCTAGC
TAATTTTTTTCTTTTATTTTGTAGAGACACGGTCTTGCCAGGTTGCCAG
GCTGGTCTAGAACCCTGGGCGGACGTGATCCGCCTGCCTCGGCCTCCCA
AAGTGCTGGGATTACAGGCGTGAGCCACTGCTCCCGGCCTTGGGTGCAAA
TTTGAGCTTTCTCACTTATTAGTGTAAGACATACAGCTAATTTCTAAATC
TTCCAAACCTCAGATTTTTTCATCCATGAAGTGAGGATTATTATAGAGCTC
ACTAAATACATGGCTTCCAAAAATATATAATGCCAAATTGAGATCAAAAT
AATAAATCTATATTACATGGGAGATCTTAATGTACCTCTTATATTATTGA
TAGACTAAGATGATCAAAAAATAGAAAGAGAGCAGTAAGGAGAGCAAGC
ATTTAATCAATAGGACCAATACATTTTAATCAATAGGATCCTCAGGAATA
TATACAGAATACCAAACCTAACAACCTGCAGAAAAACATGCCAAACATTTAG
GTACAGACATTGTTGGAAAATGCAATCTTGAAACGAGTGGACTGACATTC
AGAAGATATTAATAAGAGCACTAATGATGGGGATTGCAACCATGTCTTTA
CTGACTTCCAGAAGCTTCTTACAGTAACATGAAATCACATAATTTCTTC
CACTTTCCTTACTGTTTCTTGTCTGGGCTCTGTCTGCTTACTGTCTAAT
ATCTTGGCCCTTAAAGTTGCTAATCTTCCAAACCTCATTCTGTGACT
GGGCGCTGGTCTTGTTCATGGGCCTTGAAATACTGACTGTACACTTA
TCTGGAGCATCCAGTGCCTACCACCTGACCCAGATTCTCTATTGCGCTCC
TCCCTCTCCACCTATTGGAATTTGCTCATACCCGTGTGAGACCCCTCCC
TTTCCCCCATCTGAATTTTATCAAGACAACGCACTGCCATACTCCCTC
GTACCTGCTCTGGGCATCAGACTGAATGTTTGTTCATTGAGGATCTG
CAGCTGCATCAGTTTCCCCAGCACCGTCCAACCCCTTGAGCATGGCTAGT
CCTAAAGCAGAGAATTAGCCTTTCTATCCCTGCTGCTATACATGCTGGGA
CAAATAATAAGAAATGACAGCATTTTATGATAATGCAGGCTGCAGGAGGC
AGGAGGCAGGAATCAAATTCGTGCTTATCAAATAGTGCTCCAATTCTTTG
AATATTGACTATAGAATATGTATGGATCTATGCTCAGGTGGGTTCCTT
ATTACTCACTCCACTGAGGCCAGGTTGTGGGATTAGCTGTCCAAGAGGGA
GTTTCAGTCTCACAGCATAGGGTCATTCTGAGAATTACTGGCCCACTT
GTGTGGAGACCTCCAGAGAACAAGATCTGGGTTGGTGCCATGTACTTCA
GGAGGAGAGAAGTGGCAGGATGCCCAGCCCCACAATCAGAGGGGAAGGGG
CAGAGCCACATGTATGAAGATCCTCTCCCCAGTACGTGCCAATCACAGGG
CTTCTAGCTTTTGGGCCAAGGAAACAATGTGGGAAGCAAAAAAGGACAA
TTTTCTCTCTCTTTGTCATGAAGACTGAGCAGTTTACCAGATTCCCAGG
GAAACACCCCTTCCACTCTGGGTTGAATGTGAGTGAGAGACATTAGCTGG
AACACTAGAAAACTATTTCTTGAGCCACTCACCTTTAGCCCTAGAAAGT
GTTGGATTTGTCTTTCATCTTTGCCACAGTAGAGACTGCTGATAGCATCA
GAACCTTGGGCTCTGGAATTAGACAGATATGGGTACAAATCTGAGCTCTCT
CACTTATTAGTGTGGGATGTAGAGCACTTTTAAATCCTTCCAAACCTC
AGACTTCTCATGCATGATGTGAGGATTGTAATAGGGCCACCTAATAGGG
GTTTTTGGAGAATTAAAAAAGTTATTCAATGACAGCATTTAGCAAGATGC
CTGACCATTGAGAAAAATAACAAATTGTTTATTATTATTGTTATTATTAA
CATCTTCTGACCTTCTGACTGGGGGCATCGTATCATCAGAAATACTT
AGGATGGGATGGATTCTGTCATGGGCTGAGTCAAGGGTGCAATAATGGAG
GAGTGAAGAAGGAAGAAATGGAGGCAGAAATCCCAGGAGCCCAGCATGG
TACAAGGCTGAGCTAGTGTGTCAGAGCCTCCTTGAACAGCCACAGAGCT
TGTCATCTGGCCCTGGGAGGAACCTCTTCTAGCTGGCAGGACCAGCCACAA
CAGTGGCCAGGGGATTTCCCAGGGCGTGGGCTCCTAGGAGTTTCAATTTGGA
CCAAGCCTGCCTGGAGAGGGGTTATAACAGGGATCCTTCCCTACTGGCAG
GTGATTTACCCCTCGGTGAGAAGCTCAGGCATTTGTTTGATGGAAGGTGG
AAGGCCCTGTGCTGGGCCAGTGACTATCAGGGATGGGCGGGTGGCTGGAA
AATAGCAAAATAAGACAATATGATAACACAGTTAACCACCACACTATGTGA
AGCTACAATATGGGTATCTGTAATAGACAATTCCAATGTAGAGAATAATT
CTAAGGTGTCTTCTCCCCGCAATGCCATAAGCACACGGCCTCTGCCTG
GGTTCTCACTGTGGAATGTCTCTCTGCTCTCTCATGCCAGAGAGTGG

FIG. 3 (46 of 52)

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GAAGTACTGCTACTTT. .CACCGGCTTTCTCTGTCTCTCCCTGCAGC...
CCTCAGCCCCCTCTGCACAGGGAGGTTTCTCTCTGCTGCTGAGTGCTT
TGTACTTGTAGTGGTACCTGCACACAGGTATTGGTGTCTTGTCTCACC
ACCTTACATCACTGTAAGCTCCCCAGGAGCAGGCTTCTGTTTGACTCAC
CTGTGATCCTCCACCTCCCACTCTGTAGTGCCTCAAGCATTGAGGACAA
CACTGGCTGCCCTTAACCCAGAAATGCTGCCGAGACAGGAGGCCATGGC
CCAAGTTCCTGGAATGGGGTATTACTATGTGAGCACAAAGGCCTTTGCAC
AAATGAAGGCTTTAAAAATGCAGTCCTAGTCAGGTGGAGGAGGGCTTATA
GGATTCCCAGGAATCTGGATCATTCTCTTGAGAGCTTTCCCTTGTCTCTG
TTAAAACTCACATCCTACGGCCCCAAATAACAACAAAAATGGATGTAAAT
TCTTGAAATAACTTGTGGATGGGGGAACAAGGCCCAACCCCCAGATCTGC
CAGAAGCTTCAGGTGAGGGTCCCAATGCCAAAAAGTCTGGTATCAGAGA
GGATGGCCAGTGACCTGGGGACACATGCCCTTGTGTGTCACTCAAGGA
GCAGCAGCCTCGGCCCCGCACAGTGACCAGGACCTGGCTTCCCACGCTG
GGCAGGAGCTGGTGTCTGATGAAGGGAATGCCTGGCAGCAGTGTCTGTCT
GTCTCCTCGTGTGAGCTTACCTGGCTTGTCTGGAAGAGGCCACTCGCAT
TTCTCAATTTTTTATATTTTTTAAATTTTTTAAATTTTTTATTTTATTTT
TATTTTATTTTATTTATTTTATTTTAAATTTTTTAAATTTTTTAAATTA
TGCTTTAAGTTTTAGGGTACATGTGCACATTGTGCAGGTAGTTACATAC
GCATACATGCGCCATGCTGGTGGCTGCACCCACTAACTCGTCATCTAGC
ATTAGGTATATCTCCAGTGTCTATCCCTCCCCCTCCCCCAACCCACAA
CAGTCCCCAGAATGTGATGTTCCCTTCTCTGTGTCCATGTGATCTCATTG
AATTTCTTTAAAGGTGGAATCTCTCAGTGGGGTCTAATCTGTTGAGAAAT
ATCAAAAGAGTATCCTTGGGAATGACTGGAATTCAGAGTCATCTGGTAA
TCCTCATAAACAACCTCTGGATGTCTCTCAGCACATCTCCACCTTGAA
CGCAGGAGGCTGGTTCAAATGGAGGAGCATCGCTCTACTGCACTTTTTTT
TTTTTTTGGCCTAAAGTGCAAAAGGGGATACGTTTCATGTAAATAAATCA
ACTGCAAAATCGCTAGTTATGCTGAGCCCTGTCCCGTGTGTGGACACAAA
GGAACCAAGGCTTTTCTCCCCGCCCAACACACACATAACACACACACAA
AATCATAAAAACATACATACCCCCAACACATAACACACACACACACAC
ACAAAATATATACACACACACACACCAAAACATGCCCAAAACCTGTGTC
CAGAGATAGATCCTACTGGTGGGTTTGTGGTCTCGCTGACTTCAAGAATG
AAGCCGTGGACCTTCGCAGTGAGTGTTACAGCTCTTAAAGATGGCATGGA
TCCAAAGATGAGCAGTAGCAACGTTTACTGTGAAGAGCAAAAGGACAAA
GCTTCCACAACCCAGAAAGGGGACCCAGCAGGGTTGCTGGTTGGGGTGGC
CAGCTTTTACTTCTTTTGGCCCCCTCCCATGTTCTGTTTCCATCCTATCA
GAGTGGCCCTTTTTTCAATCCTCCTGTGATTGGCTACTTTTAGAATCCTG
CTGATTGGTGCATTTTACAGAGTGCTGATTGGTGGCTTTTACAATCCCT
TGTAAGACAGAAAAGTTCCTGATTGGTGTGTTTTACAATCCTCTTGTAAG
ACAGAAAAGTTCCTCAAGTCCCCACTGGACCCAGGAAGTCCACGTGGCCT
CACCTTTCAACTCCATAATGGCATGAAAATACATATGTTGTACAAAACAT
ACATACACAAAGTATACATGCATCTCCCCAAATATACATACACAGAA
ACATACACACAGGAACCTCAGCTACCTGTCAAAAGTCTGCATGGTGAATTGC
CTCTGCAGTGAGTAGTTAGAAAAGTGAATTTGTTTTTCAATAAATTGGAG
TCCTTAAAAATCGTTGTAAGATAGAAAATTTTTTAAAGTATATAAAATAA
AATATGTATGTCTTTGGTCTAGCATTTACACATGTAGGAATTTATCCTA
GTGGAGTAATCAATGATATATGCAAAGATTTGGACAAGCATATTAAGCAC
AGAATTATGTATGCATATGTGTGTATATATATATATATCTCATACATA
TAATAATGTAAAAGTGAATAAATCAGATGTTCAAAATTGAGGATTAGT
TAGACTATGATCTGTCCATATGTGACATACAAGTTAGCTGCCCCTTATTC
TCTCGAGCTTCAACCTCCTATAAACAGTGTCCCTTGTATATCAGTATTGG
TACAGATAATCGAACTTATTGAGGTTTTACATGGGGCAATAAAGGCAAGA
GTTTATGAATACTCCATACTACACTAGGTAGCACCCCTATTAAAGACAA
ACTCTTCTCTCTCATTTCCCTTCCCTTCCGGAACCACTTGGTTGAATCTC
TACAAGTCTCTATTGCAACTGCCTCAACATGGCACCCCTCCCTGCATCTCC
ATCTTCCCTGTCTGAGAGCAATGGCCTGCTGCCCCCACTCACATCCT
CATTTCATTCCAGAAGTGAGCACCACAGAAGTGCTTACAGTTACCCCAACC
ACCTTCTTAGAAGATAAGTTAGTGTGTTTTGTTTTGACTTTTTAAATTTTTA
CTTCTCTTTTTCTTCACAATCTCATCCCATCCCAAGAGGTTTATCAAGA
AGTTCTCTAAAGATATGTGTCTCCTTATGGAATTTAACAGAAATCAGGGA

FIG. 3 (47 of 52)

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...GATTCAGCCATCAGGGAATAACATTTTTCCAGGTCTTTAGAC
ATAATGGAATACCTTGACGTAATTAGATACACTATTGTAGAAAAGTATTG
ATGAAATGGAACGATGTTTGAGATATCATATTGAGTAGAAAAGGCAAGAT
ACATTAAGTAGGAAATGTATCTTACAAAATAATTTGTGAGACACACTCCT
ATATTTGTATGTTATATAAATGCGTATGTGAAGAAAGGCTAGAGGATGAG
ACCCACAGTCTTCGGTGAAGTTTAAGAGATGATGCTGCAGCATGCTCAGAA
AGGCTTGGTATAGTTTTTTCCAGTAATTAAGGACTGATCTTAGGTAAATT
GTCCATCCTCTCTAAACTGCACCACCTTTGTCTGTAAAACAGGAAGGAT
GGTATTTACCCCCAGGGTCATCAAAGGATTTGGTTGGAGAAAAATAAATA
AATGGGCTGAGCCAGACCTGGCACAGTGAGAGCACAGTGGTTGACTATT
GTGCTGGCCTGTTGTTCTGTGTTATTGACATGCTGCTGGTGGTGGTCCA
GAAGCTATTACTTAATTGGTTATGTGGATTTCCCTCATACTGAGCAGC
TGTGTGTGGTGTGTTAAAAACATAGCCATACACAGTAACTGACAAGGGCAA
ATGTGATGGAATAATGCAAGGAAGTGCAGATAAATAGCTAATGGGCTGTA
GAAGGAAGCTAGTCTTGGAGGGCTTGATCAAGGAAGGTCTTTTGCATG
TCACCTTTGAAGAAGAGGGGACATAGAAGAGGTATAGTGCATCCCGAGT
GTACCTGGAAGGGAACATGAAAAGAGGACATTTTTCTCTGGGACATGGGG
ACTCCACTTGACATGAACCTCTGGAATTGGGGCAAAGAACCATCATGAGAAC
AAGGGCTTCCTTGAACCTCCAGGCTCATTGGCTGATCTAAACCCTGTGT
CCCCTCTTTCCTTCACTCTCCTCTGTTTTCTATACCTGTATTATTGGACT
GGACTGGAAGCCACCTGATCTATCACAAGTACCTTGAATGTGTTGAATA
GGTGTGGCACAGTCTTAGCAGAGTGGCACTACCCCCACAGGAATTTGTT
TATACCTTTGGCATGGAATAATAGCAGGAAATGAGTGATCACTGATAACTG
AGGATGCTATTTATTATTGGCCAAAGGAATACTTGTGTTGTATTTGCATA
ACCACTCACAACCTGTTGATTACAAATGAGTACCAGACCTAGCTCCTTCA
AGTAAAGGATCTTGAGAACTGAAGGCAAACAGAGCTCCAGGAGTCCAAGA
CAGAGCCACAGACCAGGATCCCTGGCCAGGTAGGTGGTCTCTCTGC
ACTGGCTTTCAAGGCCAACAGGATGGATGGGAAGTAGAGTAGCATCTGG
CCATCTAGACCTTGCTTTTTATCCCCACTGGAAGCACATCTGAATTTCT
AAATATGATCTCTGAGACCTGCCAGAACACCTTGCTCTCAGCCCCAGTA
GCAGCCTGCTCTCTCCAGGAGGGCTTCCACTAACAAAGTAGGGCATTGCT
GGAGGGCCAGGCAGACACTAGCTTAGGAAATCCACCAACCCTGGAAATGC
TAGTCCCTTCTCTGAAGGCTCAGAAGACTGACTTTAGAGTCTAGAAAATA
TTGGTCTCTGGGAACAGATTTTGAGTGCAAAGAGATGGACTTCAGATGGC
CAGATGCACTGCTTCTTTAGGGAATCTGTGAAAGCTCCCTGCATTTATC
TTAATACAGGCAGCAGATTTTATGAGTACCCCCGAGGGATGGCCCCAGGT
CCTCCAGCCTGTGAGCATCCTTCTGTCTTCCAGCAGCACCACAGTATCTT
TATATGCTTTGGATACCTACGTTTCTGCCAGACATCTCTTGCTCTGATG
TTCTGGCTGCCAAATCTCTGTCAAGCGCCTCCAATTTTTTGTGTCCTTT
GATTTACCCCAACATGACAAAGGCAGTTGTGCTTCATGTATTAGGGATA
CTGCCAAACCACAAACAGGTTAAATCAAATAGCAGATATCCCTGTTCT
AAAGACCCATCAGCTCTACCCACCTGCTCCTGCTCACCCTCCTTATTGTT
GAGTCTCTGAAGCCCTTCTTGTCAATTTTTATTGTCATGAACAATTTA
GTTCCCTTTGTCTCACTCCTAAACCTTTCTCAAAGGATTGGATTTGTACA
CAAACCTGCCTATCTCTGCAATCTTAGAAGTGATATGATTCTGAACAAATC
ACTTAACTTTTGAATTTTTATTGGTAAGATGGGAATACCAATTTTTGCTC
CACTTCTGTCTATGTTGGCCTGGGCTGATGTTGAAAGCTCTCGGTCAAC
TGAGATAGGGTGTGCAGAAATTTATATATATAAATATATCTCCTCCAACCC
CTCCAATGAAGCAAGTCACGTGAGTCAATCTACCTAAGATATTAGGG
ATTGAGCCTCCTGGGACATTTGGTGGCTTAGGTTTTCATGAAAAGAGGTT
GCAGAGCAACTGCTTTTTGTTAGGCAAAGATTAGGCTACTGCAGAGACTC
AGCAAACCTCTATAGAAGGTGTCAGATGGTAAGTATTTAGGCTTTGCTT
GCCAGATGATCTCTCAACTAGTTAACCATGCTATTGTAGCCTCGAAGCAG
CCAGAGACGATCTGTAAACAAGAGCATGTAGTGTGGCATAAATATAGTA
CCGCG
>Contig43
GCAATAAGTCTATTTACTGTAAAGTTAATCAAATTTACATTTTCAGAACAC
TTAATCTGCAAGAGTCTTTTCCAAGACCCTATACCTAATTTTGTGTTAC
AATTTTATATTTGTTTTCTTAAAGAAGACCACCAATATAAACTATATCCA
GCCTTCATGATAAGTACATAAGAACTATGCAATAAGGGGGAAAAA

FIG. 3 (48 of 52)

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CAAAGAAAAATACCTAC TACTAATGGTTCACCTCTGAATAGCACAT...
TCATAATGATACAAGCACTCATTACTAGTCTAGGAAAATGAAGATATAAT
TGCAATTAGGAAGATCAAGAGGTAGGAAATGTGGATGTGTGGTATAGAC
TAGGGCAGGACAAAGAACCTAAATCCTCATTCTTCTAAAGATAATTGTTAA
TACGTAAAACTCAAATTCAAGAAGTAACAGTAAAAGCGGTCAATTAAGAA
ACAAGCACTAAACACCAGATAGGAAGCGAGAGATGGGGGAAGAGGGCAAG
AATCTGATTATTTTTTGCAACAAATTTTGTAAAACCATTTGACTGTTTAC
ATGTAGAACTTGGATCTTTTTTAAAAAACACAAAATAATAACTATTAT
TTTTTAACCTGGATTTTTTGAAAAAGAAGATAAAAGTCTCATTTTAGTAATT
AAAACCTCATTCCAGGTTAGTCCACTCAAACTTATATTCGAAAAATTAAAA
CTTTGGGAGGCTTGAGGCAGGCAGATCACCTGAGGTTGGGAGTTCGAGACC
AGCCTGACCAACACGGAGAAAACCCCGTCTCTACTAAAAATACAAAATTAG
CTGGGCGTTGTGCATGCCCTGTAATCCAGCTACTCGGGAGGCTGAGGCAG
GAGAATTGCTTGAACCCGGGAGGCAGAGGTTGCAGTGAGCCGAGATCACA
CCATTGCACTCCAGCCTGGGCAACAAGAGTGAACTCCATCTCAAAAAAA
AAAAAAAAAAAAAATTAAACCTCTGGAAGTTGAGTTTGCAGATATTCA
TATGCTCATTTTTAACTTGTATGTTTGGAAAATGTCATGATGAGAATTGA
GGTTGGGGGATGAGAAAAAAGAAAAACATCAACCCACAGCCCATTCAA
TTTTTCAGCCCGACCCACAGCTCCGGGGAAGGGCAGCAGGTCCATCCTTCA
CTCTTTCTTCACTCTTTCCCTCCTTCTGGCTCTTCCACCTCTAAGTTG
GAGCCCAAGAAGAGGCACTGGGAAATGGAAGTCTTTTGTACGTGGTAC
TTGCCGGGGAAGCTGCCATGAAGACCTGGCCCCACGGTGGGGAGGGAATG
CCCAGCTGAGGCCTCGTGCCCATGCTAGGATAGACTCGTCCAGACATGTC
AGGTGGTCTGACAGGGCAAGCAGCAGGAAGTCATGTATGAGTATGAACTG
ATCTGTATGCAAGGGCGGGGAGAACACGCGGAGGAATGGGGCGTGAGAAA
ACAGCACAGTACGTTTCTTTAGCAGCTGTCTCTGCTCAGCCATGGGAGTC
ACCAGAGAAAGAGGCTTGGAGGCGTTATTTTCACTGTGAGATGTGAGTGT
AAAAAAGTGCCCAAGACACAGTGAGTACCAGGGAGATGCCCTCTTTCCCT
ACCCGAATGCAGAATGGCCACAGGCCTTAAACACACACATGGTTTCTCA
GAGGAGAGAGGCCTCCACAGTGGACACCCGCATTCTCCCTGGTCAGCAG
CAGCAGGGCGAGTGCTGGGCCATCATGAAGCTTACAGGCAATGAGCTCT
CAGCAATAACAGGAACAGTGCCCTGGGGGACTGTAGCTGCAAGACCGATTT
TCATGTAAAGATGGCCTCTGAGGACTCCGAGATACACCAGGCTGAGACTAG
CTGGCAGCTCCAAGTTCTTGGTCAGAAGAGAACAGGAAGTGGGAAATTG
GAATTACTGTTACTACAATTCTTTTACATCCGCACAACCATGAGGTCCAG
AGAGTCTCTCTTATTTTTTTTTTAAAGACAGGGTCTCACTCTGTGCCCCA
GCCTAGAGTGCACTGGTGTGATCATGGTTCAGTACAGTCTTCACTCTCCA
GGCTCAAGTGACCCCTCTGCTCAGCCTCTCAAGTGGCTGGGACAGCAGT
TGCATGCTACCAGGCCTGGCTTTTTTTTTTTTTTTTTTTTTTTTTTTTT
TCGGTAGAGACTGGGTCTCTGTATTGCCAGGCTAGTCTCGAACTCCT
GGGCTCAAGTGATCCTCTGGCCTCAGCCTCCCAAAGTGTGGAATTACAG
GCATGAGACACTGCACCCAGCCAGTATAGTCTTTTAAACAGCTTTATTGAG
GTACGGCTAACATTGAAAAAACTACACAAATGTAAAGTATGCAATTTGAT
AATTTTGACAAATGTACACACCAGTGAACTATCACTACAGTCAAAATAA
TGAACATATCCATCACTCCCAATTTCTCAGCCCCCTTGGTAACCCCTCT
CTCCCAACTCCCTGCCCCCTAACATCAGACACTACTGATGCATTCTGTC
TCCATAGGCTCATTTACATTTTCTAGAATTTTACATAAATAAAATGACAG
AGTATATACTCCTTCATGTATGGCTTCTTTCAGCCCAATTATGTCAAGAT
TCATGCTTATGGCTGTGCGTATCCTTAGCCCATCTCTTTGTCTTGCTGAG
TAGGATACCATTGCATAGACAGACCACAGCTTGCTCATCCATTCACTCTT
GACAACGTTGAATTGTCTCTGTTTTTTGCAATGACAAATAAGGTTGCTAT
GTACATTCTGTATAGACATTTGTAAAAGCACAGCATTTCACTTCTCTTG
GGTAAAGACCTAAAAGTGGAAAGGCTGAGTCATATGGTAAATATATATGT
CTAACTTTTTAAGAACTGTCAAACCTGTACCCAAAGGGATTGTACAATT
TTACATCCCCACCAGCAGTGATGAAAATCCCGTACTTCCACATCCTCA
CCAAATATAGGTGTGGTCAATCTTTTTAATTTTGGACATGNTAATGAGTG
CAAAATGAGGCCCAGAGTGTCTGAAGTTACATTTGTATCCTTTTTTGGCAT
CCAAAACAGGTGTCAAGCATAGAAAAAACACTTGTTCCTTGAATGGTCAG
TCATTTACAAGTGAATTCATTACAAACCGGTAGTTCTACTGGGTAAAC
TATGCCTTACTGTCAACAGGCACATACACATACAGACAGACAGGAAGGCA

FIG. 3 (49 of 52)

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CAGAGACAAGGCAGAGC...TGATAAGAAGGTGACCTGGGCTCTAGCTC2
GCCTATCACCTAGTAAAAATATTAGTTAAGTAGCCATGAGTAACTCACTTA
ACTTACCACAGGCTCCATTTTCTTATCTGTAAAAATAGGAACATTGAAACA
GCTAATCCCCAAGGTTTGTGGATAATCAGAATTACAAAGATCAATGACAT
TTCTATGAGAGAAACATATTTCCAAGTATTTGATGGAGTACATCAGACAC
AAAGGAAAGGAAACTGAATATTTTGGAGTTTATTTTACCAAGAAA
TTACATTTTGTAAATTTTTCAGAACTACCTCCTGAGGAAAGTGTAGCTG
CACTCATTTAGAAATGATAGAAAACATCAATCTGTCTGATTCCAAAGCCAA
GTTCTTGTCTACAACGAGAAATGAAACAACTGGATCCCTACAGATGCAGAG
ACCTGGGCCCCACAAATGTGAATTCTGTTCCCTACCGAATAGAGTTACA
GTTCCATAATACAGTACTCCCTCACTTTTCCACAGTCTCACATTCCACAG
TTTCAGTTACCCACAGTCAACTGCAATCCAAAAATATTAATGAAAAATTC
CAAAAAATAAACAAATTCAGAAGTTTAAATTTGTGCTCCATTCTGAGTAGCG
TGATAAAATCTTGTGCCACCATCCCACCTGTCCAGCTTATCGTTAGTCAT
TGACATCGTCTGCTCCTGACATCCAACCATTGACATCATCATGACTCTAT
GATCCAGGATCACCGAAGCAGATGACCTCCTTCTGACATATCATCAGGC
CAATATCAGCCTAAACACTGCATCACTATGCCACATCAGTCACCTCACT
TCATCTCATCAAGGAGGCAATGGATCACCTCACATCATCACAAGAAGAAG
AGTGGGTATAGAACAATAAGATAATTTGGGGCAGGCATGGTGGCTCAGC
CTTGTAATCCCAATACTTTGGGAGGCCAAGGCAGGAGGATCCCTTGGGCC
CAGGCATTTCAAAACCAGCCTGGGAAACATAGTGAGACCTCCTCTCTCTGC
AAAAAAAATAAACAAAAATATCCAGATACAGTGGTGCATGCCTGTGGTC
CCAGCTACTCAGGAGGCTAAAGTGGGAGGATCACTTGGTCCCAGGAGGTC
GAGGCAGCAGTAAGCTGTGATCGTGCCACTGCACTCCAGCCTGGGCAATA
AAGTGAGACCTGTCTCAAAAAAAAAGGTAATTTTGAGAAAGAGACCAC
ATTCATACAACCTTTATTATAGTATATTGTTAGAAATTGTTCTATTTTCACT
ACTTATTGTTGTTAATTTCTTTCTTGCCTAATTTTTTTTTTTTTTTTG
AGTCGGAGTTTCACTCTTGTGCCCAGGCTGTAGTGCAATGAGACGATCT
CAGCTCAGGCAAAATCCCGCCTCCCGGGTTCAAGTGATTCTCCTGCCTCA
GCCTCCCGAGTAGCTGGGATTACAGGCGCCTGCCACCATGCCAGCTAAT
TTTGATTTTTTAGTAGAGGCGGGGTTTCTCCATGTTGGTCAGGCTGGTCT
CGAACTCCTGACCTCAGGTGAGGCCTCAGCCTCCTAAAGTGCTGGGATTA
CAGGCTTGAGCCACTGCGCCTGGCCTCTTGCCTAATTTATAAATTAAC
ATTGTCACAGGCATGTATTAATTTATAGGAAAATCATAGACATATAGAGT
TGGGTACTATCCACAGTTTCAGGCATTCACTGAGGGGCTTGGAACACGCC
CTCCTCAGATGAGGGGGACTACTGTCTCTCTCAATCATTCTTGATTC
AATCCTCAACACAAATGGTTTGGCCAGGTCTTGCTCTGGAGACAAATTT
GCTAAGGATTTAGAGGGGAAAAAATGTAGTTCACTGGGAAAGTCACCTCT
GCTCCACTGGACAGCAACTTAAACCCAGGCCATGACAAGTAGAAAGGCC
ACCCCTCACTCTCTTACACCTGGAGTATTCAGGAGTCAATCATATTTCA
GGACCACCAGGAGCAAACTGGGAAAACTGAGCTGCCTTGAGGAAAGCAA
TCAGCTCCACAAGGGGCTTAAGAAACAAGCTCTGGGAGGAGTGGTTGGAG
AAGAGTTGGGGACACATCAGAAATGCCATCAAAATTTCTAAGGGCTACCTC
GTGGTGTGAGACCTGTGCATCTTCAAGGACATAAACAGATGGGATAAGCA
GATGAGATTCACAGAGGACATCAAAATATTGGCTCCCCAGAAAGGGAGAAC
ATTCTAGTAACAGAGCTGCCAGCTGCAGAGTGGACTGTTTCACAAAGCA
ACAGGTGCCCTGCTCTTGAATCACCATCTTACAGGAATGCAGTAGAAG
GGACTTAACTCCTGCCCTGAAGAAAAGGTTAGGCTAGGGAACAGCTCCA
AAATTTTTTAAAGGAAGCAACATAGGCATCTACTGGGAGTTTCTAAAG
CCTTTGTTAATGAACTAAAGAGCTGGGACAGGAAATGCCAAATTAAT
TAATAGAGCCTTGCTTTAAGACAATGCAAGTGGATGGTAATGAAGGAATG
AGTCTTAGGCCTTGGATCAACCGTATTAAGCAATGCTGAGCATGGAGCCA
ATTCTGTTCACTAGATTTGCTCAGAAAGGGCCAGACGAGAAGGATTTTC
TAAAGGCACCTACTACCAAAAAGCTGCCAAGGCGTCCAATGGAGCCGAGA
GAGAATATGCTAACAAATAAAAAGTTGAACACCCTCAATAAAAAAGGGTAA
AAGTAATTAATAGAAAATTACTGAAAGCTTTTTTGAACCAAAAGTAGTC
AGCATTGGTAAAGTCTACAAAAGTGGACACTTTCATATAATGTTGGCAG
GAGGGTAAAAAGACATAACCTTTTGGAGGACAATTTGGCAACAGAGTAC
CAAAAACCTTACAATTGAAGAGAACTTTGGCCTGAGTGCAGTGGCTCACA
CCTGTAATGCCAACACTTTGGAAGGCCAAGGTGGGAGGATTGCTTGAGCC

FIG. 3 (50 of 52)

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CAAAAGTTTGGAGACCAGCTGGGGTAACACAGTAAGACCTCGTCTCTATG
AAAAATAAGAAAAGTTAGCTGGGCATGGTGGCATGTGCCTGTGGTCCCAA
CTACTTGAGAGACTGAGGCAGGAGGATCGCTTGAGCCTCGGAGGTCAAGG
CTGCTGTGAGCCATGTTTCATGCGACTGTTCTCCAGTCTGGGTGACAGAAT
GAGACCTGTCTCACCAGAAAAACAAGGCAAGAGAGAGAGAGAGAGAGAA
GGAGAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGATGGAAGGAAGGAAA
GAGAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAA
AAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAA
AGGAAGGAAAAGAAAAGAAAAGCAAGCAAGCAGGAAAAGGAAGGAAGGAAG
GGAAGGAAGGAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAA
GAAAGAAAAGAAAAGAAAAGAAAAGAAAAGAAAAGGAGAGGAAAAGGAAA
AGAAAAGGACAAAGAAAAGACCTTTGAACCCCTGAATTTCACTTTTAGAGA
TTCATCTTAAGGAAATTCATTCCAATAGAAATTTATCCCCAGGATTATCT
AAATATTTGCTTTTATTTCTTCTAGTAATTTTATGGTTTAACTTTCTCA
TGTTTAAGCCTTTAATTTATTTGGAATTTATTTTGGTATGAGAAAGTGTG
ACCTTTTTTTTGTTTTACTTTAAAAAAAATGTATTACGATTATTTTAG
AGACAGGGTCTTGCTCTGTCAACCAGGCTAGAGTGCAGTGGTGTGATCAT
AGCTCACTGCAGCCTTGAACCTCTGGCCTCAAGCAATCTCCCTCTTCAA
CTTAGGAGTAGCTGGGACCACAGGCGATGTACCACCATGCCCACTAATTT
TTTTTATTTTTGTAGAGACAGAGTCTTGCTTGTGGCCAGTCTTGCAAT
GTTGTCTCAAACCTCTGGGCTCAAGTGATCCTGTGCCCCAGCCTCCCAA
AGCACTGGGATTACAGTGTGAGCCACTGCGCCAGCTGCCTTTTTATTT
TTTAATTTTTTTCAGATGCTTTGTTGGTTCCAAAATAGCACTTATTAACCCA
CGCTTTCCCTCTGGTTTTAAATACTGCAAGTTTGGCTTTGAAATACAA
CCCACTGCCTTATTCAGGCTACATTCAGGAAATCTGAGACCAAGAGTCT
GAAGGCCAGTTTCTTCTCAAACCCAGGAGGTGGTAAATGTGTCACTT
CCACACTTTCTATCTATTTCTAAGAACTCCTTCTTTCCAACTCTGACAT
GCCCCCTGGCTCAGGTCTATAGAAATCCCAGGGTCCACAGACAAAGCAGA
ACTCACTTATGGGGAATCTGGGAAATCTTATCTGTTAAACCTGCCCCA
TATGGTGACTCAGATTGTCTAAAGCCCAAAGCATCATTTTCCACCCCAA
CCATTTCTCTCTCAGACTTCTCTATTTCTGTGGTCCAGAGTCAAGATCT
TGATATTACCCTAGAGTCCCCCTTCTGCTCTCTGCATACCAGATGCCC
CTCCCTCCCAGATCCATTCTCCACCTCCCTCCCATCAGTTTGGTGGG
CCCATCACCGCTTCCCCTGGCCAGGCTCTCCTTTTGTGCGCTTGGAGCA
GCAGACTGATCTCCAGCCTTCACTCACTTCATGTGGTAATCTGTTGTGT
TCATCACTGTGAGAACTCTCTGCTATCCCTCACTACTCTGCTGAAAACAC
TCTAGTGGTCTCTCATTGCTCATTAAATGAAAGTCTAGATATTAACGCTAG
AAGGCCAGCACAAATTTGCCCTATGCCACCTACCTCTCTAATCTTTTCT
CCTTACTCTGACAGACTCTCCGTCTGTCAATTTATGTATTCTTTTATTGCT
CTCTTCTACTTTTAGTATGAACTGGATTTATGGATTTTTTTTAACTTGCT
TTCAAGTATGGAATAAAGAATTTTATTTATTTATTTATTTATTTGA
GACTGGGTCTCACTCTGTTGCCAGGCCAGAAATGCAATGGTGCAGTCATA
TCTCACTGTAACCTCGAATTCCTAGGCTCAAGCCATCCTCCTGCCTCAGC
CTCCTAAGTAGCTATGACTACGGGTGTGCATCACCACATCTGGCTAATGG
AATAAAATATTACAATGCCTAATCTTAATTTTCAAAATTTTAAATTACAT
TGTACCTAATGCCCATGCATTTACTTTTTTCACTGGGTCAATAGCCCTCA
CTTTGGCAAAGGTCCCAGGCCAAGGTAAGGCCTTACTTTTTTCCAACTC
ATCTTTTGAAAGACATAAGTGCCTGTAAGTTGTACCACATTAGGTTCTAG
GAATTTTTTCAACAAAGACTTTATCAGACTATTTTCTCTAAGTTGAGAAA
GAGCTGGGGGCAGAAATATGGCACTGAATGACTGAAGAGAAGGCACTGAAA
TCAGGCCAGAGGTTGCTGGAAAGAGCAATGAGGAACACCAGCAGCAATGA
GGAGCCGGTGATGATTTTGGCTTACAGGGAGGTGTGTACCACACCGATT
TTATCTCTACGTGGATGAACCACAGCTGTGGCTCCCTTGTCTCCAGGAC
ATCACACTCTCCACATTCCTCCCATCTTCCGGCTTCTGCTTCCCGGGC
CCTCATCTGCCCCATCCTGGGTGAACACTGGTCCGTCAACTGCTGGGCGT
ACCTTCCCGCTCTGCACACCCTCCCTGGCCACCCCACTCTCACGGC
TCGCACTGCAGAGGAGCCGCATCTCTAGCTCCAGCCCATCTGCCTCTTCT
GAGCTCTAACTTCAAGCAATTTTCCCTCAGAACCATGTCTGGCTGCTCC
ATCATACTTCAAAGCAATTTTCCCTCAGAACCATGTCTGGCTGCTCC
CTCCAGAAGATACATCTCTCAAGCACATCCCCGGGCTCTCACCTGGATG

FIG. 3 (51 of 52)

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ACTGCATTACCTTCTC ACATTTGCCCTCCTTTGGATGTATATAGA.
GTTTTAAATACAAATCTGATGTGCTTGCTCTCCTGCTTGAAACACCTCA
AAACTGCCTTCAGGATAAACCACTGCCCTTGACATGTTACAGGTTGCCC
ATGGCCTGGCCCTGCCATCTCTTCAGCCTCATCTCATGCCCTTGCCCC
TCGCTCTCTGGGCTTCTGCCTCCCTAGCCCTCCTTTAGGTTCTCTAACAC
ACCATAGTCTTCTAGTGTGGGGCCTCTGCAAGTGCTGTTCCATTGCC
TGAGACATGAATCCCTCTCCCTATCTCTACCTGCACCTTCATCTGATTAA
TCCCTACCCTTCCTACTCATGATGTTGCTTTCTCAGGGACTCTCTCTGAC
TTTTTAACTAATCAGGGTCTCCCCAGTATATATCTTCATAGCACTCTGT
ATTACTCCTTTCTTAATGACCACCTGCTGTAGACTGAATGTTTGTCTTCC
TCCAAAATTCATATGTTAAACCTAGCCCCAAATGTGATAATATTTGGAG
GAAGGCTCTTTGGGAGGCAGGCCCTCATGAATGGGATTAGTAGCCTTAT
AAAAGAGACCCCTGAGGGCTCCCTTGTCCTCCCTCCACCGTGAAGGATGCA
ACAAGAAAGTATGGTCTATGATCCAAAAGCAGACCCTTGCCAGGTACCC
AATATGCTGGCACTTGAACCTCCAGCCTCCAGAACTGTGAGAAATAAAT
TTCTATTTTTTCATAAGCCACCGAGTCTATGGTATTTTGTATAGGAGCAC
AAACAGACTGATGTGCCACCCCAACCATGATTATACGTGTAATTTATGGTT
TCTCTGCTAGTAGGGATGCACCATGGGGTTAGGAACACGCTTTTCTTAT
TTCCACACAGTCTTTAGCTCTAAGCATGTTCTCTGAATCAAAGATCCCCA
TCTTTTATGAATGAAGGAGTCAGTGAATGAATTAATGAAAGAACTGATAA
CCCTCAATAATTATTCAGCCTTTTATACCTACTATTAACAAGCTTGCAT
TCTACTCCAAATTTATTTGGGCTTTAACTCTATTTTTTGGCCAGCCACATTT
GACATTCCTCTGAAGTAAATCTATGCTTTCCATCCTAAGTCAAGGAAGGAC
CTGGACTAGTAGGGCCAGAAAGGTCTAAATTCATGGGTGGGAGAGAGA
GACTAAATCTGAAAGGAAGAATAGATTGAGCAAAGGTGTAGAGATTGGGG
AAGGCTGGACATTTGGAGAGAAGGAAAAGGAAACTGACACTAAACCAAAC
AGTCTCACAAACACAATCTCATCCTTCCAAAACCTCTGTGAAGTAAGAATT
ACTATCCCAGGGCCAGGCACAGTGGCCCATGCCTGTAATCCCAGCACTTT
GGGAGGCCAAGGTGGGTGGATCACCTGAAGTCAGGAGTTCAAGACCAACC
TGATCAACATGGTGAAACCCCATCTCTACTAAAAATACAAAATAGCTGG
GCATGGTGGTGACACCTGTAATCCCAGCTACTTGGGAGGCTGAGGCAGG
AGAATCATTTGAACCTGGGAGGTGGAGGTGTCAGTGAGCAGAGATCGTGC
CACTGCACTCCAGCCTGGGTGACAGGGAGACTCCGTCTCAAAAAAAAAAA
AACAAAAAAAAAAACCAAAAAAAAAAAACAAAAACAAAGATTACTATCCCAG
TTTTGCAGATGAGGCAATGGAAGCTCTAAAAAGTTAAGTAGGAGAAACAA
ACATGAAATGTATGTCTTATGCTTTTCTCTCATCCTATTTCTCTCAGCCTGG
AATGTCCATTCTCCCTCCACTATGCAAACTCTAACTCTTCAAGCTAACACA
TAGCAATGTCTGAGAAACCGTCCCTGTGTTCACTCTGTTAGCCTCACTTG
CTCCCTCCCATCCCTCTGTTTCTTTCTGTTATAACACTTCTCTATTCT
GCTGGCATCACAGTCATCTCCACCTGCCTTCTCACAAGTTAAAAGCTTG
TTAAGGGCAAGTGGTGTCTTTGCCACCTCATTCCCCAGGGCTTCTAACA
CAGTGCCTCATGCATGACAGAGTTGTAAAACAGGTTACCAAGCTGGCTTC
AGGCAGGTTTGCATGGAACTGTGCTTTACAGGAATACCTGCTCCCCCAG
GCCCTGGGTCTTCTCTCTGAGTCCAGGCTCAGACTCTCTCATCCTGCTCG
TTCTCTCTTGGGGAGCCACAGTAACTTTGAGCAACTTTGCATGGGATAGA
ATGGCCTATTAGGGGCAGCACAAAGACCCCATGGAGGGAAGAGTACAGAA
AGGGAAAACGATAATCATATTTTTTTAAGATGTGCATTTTCTTAACAAAA
TGCTCTAGTACTTGTCCAGACTTTCAAACCTCAAAAACCTAAGCGTCTTTT
TCTTGAAGATCATCAAAGGCCCCAGTGGTCTTTCAGGTATGTCAAGCTTT
CTAGAAAATAAAGGTAAGTCATAATCACTTAACACACATGGCTAAATGGC
CATTTCTTCTAATTTATCAGCAACTGTTACATATTTCTATACTAGAAAA
AATTTATATTTATACTCAGGGTGGTAAGTTAAATTTGCCATCGAAGTAAA
GCAGAAAGAGCGTAGCATGTATGTATATGTAACCTCACTGTGCATGAGAC
AAAGATGTCTTGAGGAGAATGAGTCTAAGATGCGCCTGAGCAATAGTACC
C

FIG. 3 (52 of 52)

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>Contig1
GCACCCATGTTTCTAAAGGGCATACCAGCCATAATAACAGGATGGGTGAG
GATATAGACAGCAGATGACAGAGAGAGAGTGAAAGCTGGGAATCCCAGC
TAAAGGCATCAGGTTTATGGAATGAGTAGGGGACAATACTGTGTGTGTTT
ATACACACATGTATATGTGTGTATATGTATACATGTTTATGTATATATAT
AATTATATGGTACCATTTCTAATTGACAAAATAATCTATCACATTTTACA
TTATCAGATTTTACATCTATTGTTCTAAATACACTCAGTCATCAGCCCTG
TGTGTGGGCTCTTACCCATCCCCATGCACACCTCAGCTCAACCACTGATG
GATGGATCATCTGCCTATCAGAGGTGGCATATTCAGGTGAATCCATGGCC
ACAGCTGCAGCACTTCCTACCCACGCAGAAAGGCTCCACAAGAGGAGGCA
CACCCGCTCTGACTGTCCCTAAGCTCCTGACATCTTACCCCCATGAACT
GCTGCTCCTGGGTGCTTCTGCTTGCCTTGCCTGCCACCCTTGACTGTTCT
CACCATTGACACAGCTGGTGCCCGATGCAC

>Contig2
NAAAACGAATCGTCACTATTGAAGCCTGTCTCTCANC GGATCGTGACTAA
GAACCCCTCCTTGCTTCAAGTTGTCTGCTTCTTAGGCAGAGCCACCC
TACATCTTAAATATATTGATTGATGACTTACGTCTCCCTAAAATATATAA
AACCAAGCTGTGCTCTTACCAACTTGGGCACATGTGGTCAAGACCTCCTG
ATGCTCTTGTATGAGTGGGTGGGTGTTCTCAACCTTGAAAAATAAACT
TTCTAAATTAAGTGAACCTGGGTGAGATTTTGGGGTTCACAGCAACAA
TTTAAAAAACTCACCATTGACCTGAAATTTTGACCTTATGCTGTTGCTCA
CACTCCTCCATGAAAAATAGACGCCATCCTATGAGTTCCCTCAGCCATGTC
ATGCCACACTTCCAACATGTGTCCCATCCACCATCTGTCTTCTTATTGC
TGCATCCTACCCAGGCCCTGATCTCTGGACCCATGTTGTATAATTAAGA
ATTTGGGGCTGGGCATCGTGGCTGTGGCTCACTCCTGTGATCTCAACATT
TTGGGAAGGTGTATTAGTCAGGATTCTCCTCGAAGGATGCAACCCTAGGGA
TCCTCTCTATGACCCTATGTCTA

>Contig3
CGCGCTCAACCGACCGATTTGCGCGAACCTGCCCCATGCCCCGAGGACAGTG
TAATCCTAAAACGTCCCCTGAATCATAAGGATATGAGTGCGAAAGTACGG
TTCCCTCTGTCAACACTTTCTAACAACGCTATGTCCGATCCGTGCACTAA
CCCCGCCCAAGTCACTGAAACACTGATGGGGCGCTTCTCTACAGGTATCC
AGGGCCAATACCACTACTCCCCTCCTCCCTGTCCCCCTTCCACTCTCTAG
AGGCCGCGGATGCGATCCTCTATTAGCACAACCGAAAACGACGGTGAAAG
TACCACGAAGCTCACGATCTGATCGGTGCGCCCAATGCGGTACAAACGGCT
GTCATCCCAACCCCGTCCCATCCTCCATATTGCCCCCCCCCTATGAGGAT
GGCCCTATCATCATGACCTCCAAAATTCTGTCTCTCCCGACGTAATGCC
GCCCCCTGCAACGCTGACACCATCAAGTCNGTCACCTCCCAAAATACTCC
TCCTAATCACCAGGCCGAGTATCCCCGTTCCACAATACTCCTTGAGAC
GGGCCGATATCACACAC

>Contig4
NNGAGTTTAGGTCAACTAGTAACAAGTGGGATTTGCGACTCAGGTCTATC
TAATCCTCAAACCCACGTCTGGACCCCTACACAGACTGCCCTCCCTCAG
TCCTCTGTGTGGCCTCAAGAAGGGTCTGGACATTCAAGTTTAAAAATCCA
TCCAAAGAATCTATGGACCCAGTGGTCTCTGGAGTCAATGTTCTGAGGCT
CAGAAGGGCCAGGCAGGAGGGAGCCGCTCTACACAGTCTGAGCAGAGT
GGGCTGTGTCCCGGCACAGCAGGGGAGATCATAAACAGAATTCTGCCCTG
GGCCCTATTTAAGTAGGACCTTTAGGCTGCCGGTGTCTGACCAAGGTC
CCANGTCTGCACGATTGGCTGTGTGTGGAAAATCTTCACTCCTTGCGGCC
TTGTCTTGGCAGAGAGCACCGCTGCTTCTCTGATGGCCACCAGGGGGA
GGCGCTCCCTGGGAACGGTTTGAANGGGAGCCTCACCCACACGTCCTT
TCCGTGGTACCCAGCACAGCTGCTACCCATGGTTACCCACAGGCCCCAGC
TCTGCTCTGAAGAAGGAGGAGTGGTGGCGATCANGCCTTGTCTGCATCCC
GTGGCTGCCCTTTCTTTTCTTT

>Contig5
GGGAGCTAACCGCTCACTGGGATTACAGGTACGCACCACCACGCCTGGCT
AATTTGTATTTTATGAGAGACGGGGTTTCTCCGTGTTGGTAAGGCTGG
TCTCGAACTCCCAACCTCAGTTGATCTGCCCGCTCAGCCTCCCAAGTG
CTGGGATAACAGGTGTGAGCTACCATGCCTGGGCTTATATGTTTCTAGTC
CAACATTTAGCTACCTTTTTTTTTTTTTTGGAGACGAAGTCTCACTCTGT

FIG. 4 (1 of 61)

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TGCCCCAAGCTGGAGCACAGTGGGCACAATCGTGGCTCGCTGCAGCCTCAAC
CTCCTCAGGCTCAGGTGATTCTCCACCTCGGCCTCCCTAGTAGCTGGGA
CTACAGGTACGCCACCTACACCTGCTAATTTTTTTGTTTTGTATTTT
TTGTACAGATGGGGTTTTCTTATGTTACCCANGCTGGTCTTGAACCTCTG
GGCTCAAGCAATCTGCCTACTTCAGCCTCCCAAAGTGCTAGGATTACAAG
CATAAGCCACCATAACCGGCCTACCTACTTTTAACTTGTGGAATTTTCTA
TAAGGTCANGGATGCCTGNGGGAACAAAAGTTTTCTCCCTTGGTATATGCA
AGTAAAATCCACATGCTGCCTCCC

>Contig6

AGGACTGTAGCTGTTGTCTAGTCACCAGGCTGGACTGCTTGGCATGATCT
CAGCTCACTACAACCTCCACCTCCTGGGTTCAAGGGATTCTCCTGCTTCA
GCCTTCCAAGTAGCTGGGATTACAGGCATGCACTACCATGCCCGGCTAAT
TTTGTATTCTTAGTAGAGACGGGGTTTTCGCCATGTTGGCCAGGCTGCTCT
CAAACCTCCTGCCCTCAAGTGATCTGCCTGCCTCGGCCTCCCAAAGTGCTG
GGATTACAGGCGTGAGCCCCCGGCCACATGTAAAAGTTTATATCTCTGT
TGTTTTACCTTGTTTTTACCTAGTCTTTCAGTGATTTGAATCTTGATTC
AGTCTTTTGTATTTTAGTGGTACTTCCAGCTTTGTGTCTATCTGTGGAT
GACATATAGTCTTGCTTCTTCATGCCAATTTAAGAAGACTGAACGGGAA
TAGGTCAAAGGCATGGCCATGAGCGATTTCTCTCCAGCTTTTCATGGTGT
TCAGCTTCAAATCTATTACATATTGGACCTGCAAGCCATCATCTTATCC
ACAGGCTATCATATAGGTGAATGTAAATTGGGTTTAGGTGGCCAAGCTG
AACGTGAGATATNTTC

>Contig7

AGCATGTTCTCTAAAGGCCTATCAAAGCTGACATCAAAGGGATAAGTTCC
AGTTACCCAGCTGAAGGGAAGGAGGTGTTTCAGATAGAGGAAGGATAAG
CATGACCTATTCAAGGCCAGTGAAAGAAGCGTGCAACGGCCAAGTCAGGA
GAACCTGAAATTGTGTCAAAGAGCTTGGATGCAAAGAGCCGTGGGAGACT
ATTGGGGGTTTTAAGCAGGGATATAATATTCAATCAAGCATGCAGTAAAA
GGTCACTGGCACCTGCCATGGGCCAGGACTCGGGCTCTACATGATTGCGT
CTGTTTTTGGAAATATCACCTGGCTGTGAGATGAAGAACAGGTAGGAGGG
TCACAAAACCTGAAGCAGAGAGACTGTTGAGGAAGTAAGCTGTTTTTGTG
TGGACTGTGGCAATCACAGAGGCAGAGGATATAAATGCACAGAGACACAA
GGCATGTGGGAGGCAGAAGGAATCAAATACAATGAGTGATCAGATGTGGG
GTTAGAATGGTGAGTGANAAAGACATACTCAAGGTGACACGCCAGGTAT
CTGGGTGGATGGTAAGACATTATGGAAGTAGAATCGAAGAGGAGGTGGG
ATGGACATTCTTCCGTTTAGAGGGGTTACCAGGAGGATTGCGCGAAC
ATGGAGAGGATTAACCAGGAATCCGGTGCCTTTTTCCAAACTGGGTGGA
GGGG

>Contig8

GGTGAATGCTTTGGCACGCTGTGTAGATTTTAGGTGACGGGTGGTGACAA
TGAGTCCGTGTGAGCGCTGATTTTTTCGGCCTTTAGAGCGAGATTTATA
CAATAGAATTTGGCATGAGATTGGATTGCTTTTAGTCAGCCTCTTATAGC
CTAAAGTCTTTGAGTGACTAGATGACATATCATGTAAGTTGCTGATAGGT
TTCCAGTTTTCCGCTCCTAGGTCTGCATATTGTACTTTTCTCTTACTCG
ACTTAACCAAGTACCAACCCAGCTTCTCAACGGATTTATACCATGGCACTT
TAAAGCCAGCATCACTGACAATGAGCGGTGTGGTGTACTCGGTAGAATG
CTCGCAAGGTCCGCTAAAATTGGTCATGAGCTTTCTTTGAACATTGCTCT
GAAAACGGGAACGCTTTCTATAAAGAGTAACAGAACGACCGTGTAGTGC
GAATGAAGCTCGCCATACCATAAGTCGTTTTTGTCTCCCGAATATCAGACC
AGTCAACAAGTGTCAATGGGCTCGTATTGCCCGAACAGATTAAGCTAGCA
TGCCAACGGGATAAACGAGTCGCTCTTGGTGGAGGG

>Contig9

GGGGTGGGGCGCCTGGTGTCTTCTAAAGAGGATCTCCTGCCAGAAATGGTG
TGCTGACACTGTTGTCTCTTGGTGTGGAACCTTGGTGGGAAGAAAGGT
TGGAAAGGGAATTTGATCCTTGGATTAAACCCGAGTTGTACTGATG
CTCACAAGACTATGGGAAGGATAAAGGCAGGTGAGTCACTCTAGGATGGC
TCANTGAGCTCCACAGAGCTGGAACCAAGGCACCAGGAGGGATTAGAG
CAGGCCTCAGTGACGTCAGCTGAGTGAACCAATGAGCAGGTGATGGGTC
CAGGCAGAGCCCTGTCTCTTTAGGCAAAAACCTTGAAACACCGTTCCT
ATCCTAGCCTGTGTTCCACCCAAAGCTGGCCAGTCTCAGGCCCTGCCTG

FIG. 4 (2 of 61)

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AGCCCCAAGGAAGTGGTATGGTGAAACAGAAGGGCCATTCTGTCCAATG
TGTGAGGAACCTTCATTTCAGACTTGTGGGAAGCCCTGATGTTCAAAAACC
TCAATGATATCATTTCATTTTCCCCATCCATTCAATGCCCATCCAATGCCC
ATCCGTTCAATGCCCTTCCATTCTCTTCAGGGAAATGAAAATTGTTCA
GAAATCCTTTCTCTTCGAGAAACCAACCAAAACCAAAACCGCGAAATTCA
CTAAACTAGCCAAGACACAATCCTGGGTTATTTTCTTTTCCCAAACCTC
CTGTGTTTAAATTAATTCTACCCTGGTTCTCGGCCCTTACTGCGAAGGTG
AACTCACCTAACCTCTCCCAAACAGAGAAGAACTTCTCTTGGTAAATG
GGTTTTAACACTTCTAAAAAACCCCC

>Contig10

GCTATGGTTCTAAAGGTAATGGACTATGGCGTACACAACGTCTCGCTCAT
CGTCTGCCAGGAGGCTAAGGTATCCACGGACAATCGCTGAGCAACAGTGT
CGTTGATCCATCTCTGTACGCACTTGTCAACATGGCAGGAGTACGGGAGC
TGCGAGAATCCTCTCTGTGATGTCCACGGAGCATGCCGTGAGACAACG
CCACGAACGGCCCTCGGAGANANCTACTCTGCAATGAAGACGTACGATAC
ACACGTAGGAGTCTTAGCTCACCAGCCGTATCTAGGTATACTGTACTCGC
GGATACTCACTCGTGCATGCGGCAATAGATCGATACGCAGTCTGTCACGCC
CATGCTCTCAGTGTGTGACCTTCTGGCGGTAGCGTNGTGGGCGCTATTAC
TGTGCGCAGCAGGCGCTCGTACATGTGTGCGGTAGCGATGCCAGGAGCT
GTAACATAGCAAGTCGCCCCCTACTCCTATCACTATCCCTACGTGGAC
CGCACTCGAGATCTGAACGCACGTCTTAACCTGCCAGTACTCGTGAGACC
TACTCTGCGCAAGCCTTGGCTAGGAGATCCTGCAGCGCCGGCAAAGAATC
AGCTATGATCCCCTTGGCATTATCGCACACGCACCATAGAGTATGTGCAT
ATTAACCTCTGAATGTGCTGCAAGCAGACGGTTGCTCAACATATATATGG
ATGTGGGGAATCGCCCTGGTCACCGCACTTGGCGTCAGGAGGCACCAG
CACGTCTGAGTGTACGCACGTTACTC

>Contig11

GGCCGAATGGTGAATTCATCCGTCTCTCGAGGGGGTGAAAGACGGGGAG
TTATGCTGTAATGGCACCCTCACCCTGGGCTTATGAGCAGACCTAACCC
TCCCANAGTGCTGGGATTACAGGCATGAGCCACCCTGCCCGGCCAGTAT
CTGAACCTCTGTGGCCAGGCAGAAAAGGTCCTGTGTTACTCGTCTCCTTT
ATCATTATGTTCCATATCTCCCATTTGCTAACATTTATGTTTCTGCTCC
ACTGGATTCTTTGGATTTTCTAGAACATACCCATGCTTTGCATTGCCTT
GGTCTTTGAATATTTGGTCCACTTTTCTGCAAAGTCCCCTCTCACCTTA
TCTTCTGGTAAACTTCCAGCCAACACCTCTTTACTAACAGAGAAACAT
GGTTCAACTGTGCACAGGCTTGCACAGAACTGTTCTCATATTGTCTTGT
CATTGTCAATGTGGCAGAGATGCACCTTAGATACCTCTTTGAGAAAGGAC
TCACTGCCCAGCTGCCCTGGCACGTGATGAGCTGATAGCTCCAGCTATAGA
CTCCTTTAGGGTCAACCTCTGCTTTCCAGTTGAGATCATATCCTTTGCAG
GGTGGCCTTCCCAGTGATGACTAAGGCAGTGTTACAATGGCCTAGTCATT
TCCTCCCAATGCTGGACTCCCAATGAACCATCTGCTCCGGAGCTTCCAC
TGGGCAGTCAGAGACCTTAGCTAGTCTGCCTCCGAATCAGAAGGCTCTCT
CTTGCCACTCTGGCC

>Contig12

GCTGTGTCTAAAGATTACGGCTGTAGTTCCAACCTCCCGCCGCCCTCTAC
TGTGTCTCTTAATGGCAGTCATTACCATCTTCTGTCCCTCCCCTTCA
TTTCTTGGATGGTGAATGTCACCTTTGCTGCAACAGAACCCTGTCCCAATC
CTTGATGGTTCAATACACATAGACATTCTTTTAAACAGGGCGGCCTCT
CAGGTCTTTAATTTCTTCCCTCCAATAACCTTGTGATGATCCCCAGCT
TAGCCACTTACTGCCAGATCATTACCAGTAACTCCAGCCCCCTCTTAATT
CTAGTTTCTAATATCCTAATCTGTGACCTCACATTCCAACCTCTTCAATC
TTATCCCCTGAGTCAAAAAATCCTTTGATCCATGCAATCCATTAAAGTCAT
CTACCTTTTACCATTCTTCGCCCCACTAGGGTTCTCATTCTTTATTAC
CCATATGAAATTCCAAGGCCTGTTGGAATCACTCCCTTGCAGCCACTGTC
AATACTCTGCCCCCTTTACTTCATCACCCCTTATGTGGCAAAACCACAGC
CCTGGTGGAGTCGATCCTTACCCCTGCTCTGTGCCAACAGCCGCACACGC
ATGGCTGATGGAGGTTGGAAAAATCCACACATGCAAGTGGGCGCTGTATGT
CCATATACGTATCCAACCTCCAGCCTTGCATATGCCTCAGTGTCTGCCTGA
CAACACATTATATGTTTTCTTAGTTCTTCTCAGTCTCCTGGGTGCCTAGG
TGAGTATCTCAGACATCCTTCTCTCTGCAAAGCTCCAACACCTCCACG

FIG. 4 (3 of 61)

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TCACATTCAACTGATGACGTGTGTCTCCTATGTCACCTTAGATCACAGAGGC
ATACATAAAACAAATCCCAGCCACTGCCAGCACTCTGCACATCTGCGAGCA
TGGCACCCCCAATCTAGGCCTTTCTGCTGTCACTTGGGGTGAGCTGATT
ATACTCGATCCTAGTCATTTCTACTTATGCAC

>Contig13

CTTAAGGCCTCCCTCTAACATTTTAATTTAAGATTGAAAAAGCAAAGATT
ATTCTGTTTTGGCTGCGCCTATAGTAAAGTAACCCCTATGNCAAATTTTG
ACACCTTATAGTATTTGACAGGGATAAGTATAAAATTGCTTGATTGATAC
ATCCACACCCAAATGTATGCTGGGAATGATTTTGTTCACGGCACTCATT
ACTTAATTTTTAAACTCTTATTTAAATTTGCAATGTTTTAAATGACCAT
CACTTAAAGTAGTAATCAACAGAGGTTAGGAGAACATAACAATACTCTTT
CTCTTAGAAAATACAACAGAAATATAATTTTTTACAGTTTGTCTCCCAA
CTTTTCTCTGTAATAACATGCCTTACTCACCTTTACAATAGGTTTGTGT
GAGAATCTTGTAATGTAAACCTGGGTGTTCTGTGAAGCATTTTTAACT
TCTAGTTTACACTGACTCTTATTCAAGTGTTTTTAAAAATATATTTAAAA
AACTGGCCAGGTGCGAGTGGCTCACACCTGTAATCCAGCACTTTGGGAGG
CCAAGGCGGGCAGATCACAAGGTGAGGAGTTTGAGACCAGCCTAGCCAAC
ATAGTAAACCTCGTCTCTACTAAAAATACAAAAATTAGCTGGGCGTGGT
GGCGGGCGCCTGTAGTCCCAGCTACTCAGGAGGCTGAGGCAGAAGAATCG
CTTGAACCCGGGAGGCAGAGGTTGTGGTGAACCAAGTTTGCGCCAATGCA
CTGCCAGCCTCTGCAGNGACAGCC

>Contig14

GGGGGCGGGCCGAGTGATCCTAAAGCCCGCTCGCTTCACAACAAAGCCTA
ACAGTCCAATCACTTAATGCTGCATTTATTCCTGGGGAAGCAAGTCTCCT
TTGCACTTTACACAGTGAGATAATCAGTTTCTCATGTGGACCACTGGGCC
AGGAGGGCCTGACAAAGGGCAGTCTACATTTAGACTGGAACTGCTCCC
AGAACTATTTCTTTCTAGTTCCACCTCGGTCTGAGGTGCCTGAGGAGAG
GGACTCAACAGAGGAAGCAGGAGCATAGCTCAAAGTCTCAGAACATGGAA
GAGGAAAAGAACTCCTCACAAGATTACGTAACCTTACAGGCGTGTGCTGCT
TCAGTAGAAGTTTCATCTCCCTCAATCCTGTACACTTTCCATACATTAC
ATACTCAAACCTGGTCAGCCCTATGGAGCAATAGCAGCAAAAGTTATTCTTA
ACAGTAATTAAACAATATAAAAGATCCCATTTAAAAATGGTTACTGGTCAG
CCGGGCGTGGTNNNTCNANCCTNTAACCCCANCACTTTGGAAAGCATGCG
GGCGATCCCAAGTCTGATATCGAAACATCTGCCTAACATGTGCAACCCCT
CTCTACAAAATACAAAAAATATCCGGGCTGTGTGTGGCGCCGTATCTCA
CTACCCGGAGCTAAGTAAGAAATGCTTTACCTGGAAGCGATTFTTTTACT
TATATCCCTCTCTTACCAGGGCGCGACCAAAATCTTTAGTATAGGAAAG
TTTATTGTTTTATGCCTTTGTCAAGGCTCTACTGTATCTTTTCTGTCCAC
TCAC

>Contig15

GGTTCTGAACAACAGCAGGCGATTCTAGCCCTGTACCCGGGGCATTGTC
CAACACTCGACAGGGCTGAATTCGTCCATAACGGTGTGCCCTCTGGGAT
ATAGGATGAAATGAATTGATCTGAGTACCTGGGATGTAAAGTTACTAAAA
CGCCAGCTAGGTTACGCCCCGATGCTTAAATATGATCGTGGCCTACACC
TCGTCCAGCAGAAAAAGTACCCTTTCTTCAACACCACCTCAGGATCCTCC
AATTTAGGAGCTATAAACTCATGACTCTTTATTTACCCCTGCAGATTTC
TCAATCCAATAGTGTGTGTCTCCCTGTGAACTCACGGATATACCGATTTT
CCCCACGTCAATTTCCACACGTGCAATCGCTTAGTCATCCCTATGTATGA
GAATCATGGATGACTATGTTGAAGTCCATCTATAAAGTTCAACCCCATC
TCCGTCCCTGATTCCCCCTCCCCAAGATCACAACGCGACTCGACATATT
GTTATCGCCCAAGGGACCTCTTGCATCCCCCATATCCACTGGTCACCTCC
CCTCTTGGCTGGAAGTACCCGGGAAGTTCTCCACATGTTGT

>Contig16

TGCGAGCGATGTTCTTAACTTTAGCGCCATTGACTCGAGCATGGTCATG
GCTGTTTCCTG

>Contig17

AGGGTGTTCTTAAAGGATACTACGTTCCCTAAAGTCCAGAGAAAAAAA
AAAGTAACATAATGTGGCTTATTTGGTATAAAAAATTTACAGGAAGCATT
GTCAAATATGAAATAGTGTGTTTGGTTTTGTTGGGCTGTATTTGTATAAAT
ATGTTATTGGTATGTGTTCCAAAATTATAGGAACTCCTATAATTCTGAT

FIG. 4 (4 of 61)

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ATGACTTGGTGTACATTATCAGTAATAATTATAATTGTTATGGTAAATTA
TTGTGTGCCATGGAGGTAACAAATTTCTCATCAAGTGTGTCTTTGACTA
TGGTTGCCCTAAAACCTTTTGGCATTACAGACAATTGTCTTGCTTTGGT
CCTCTTTAGAAGGTGGTTTTATAATCAGCTATAAACTCTAACGGGTGCT
CTTGAATGCAGGCTTAAGATAGCTTTGGAGACTGTGACATCAGAATAGAG
GAAAAACTTTCAGTATTCATGGAGTGTGAAATATTCATGAATATCAAGC
AAAACAGGAATTAACCTTCATAGATGGAATAAAAGAATGCTGAAGTAATC
TTTTTGACTTTTTTCTTAAATGTTGATCCTTCGTTTTGTTTTTCAGAG
TCAAGGAAATTTTTCTGTTGAGATATTGACAGCTTTTAAACAATTAAGTAT
ACTCCAGTGAACACAATTTGGAGCATATTTGTGTCTCTCTATATATATT
GGAAACAATNTTTGAGTATTCCTTAACCTTATTGCAATATT

>Contig18

GGTTGTCTGCTATACCAGTAATGGGATTGCTGGGTCAAATGGTATTTCTG
GTTCCAGATCCTTGAGGAATTGCCACACTGTCTTCCACAATGGTTGAACT
AACTGACACTCCCACCAACAGTGTAAGCAATTCCTATTTCTCCACATCC
TCTCCAGCATCTGTTGTTTCTGACTTTTTTAATAATCGCCATTCTAACTG
GCATGAGATGGTATCTCATTGTGGTTTCAATTTGCATTTCTCTAATGACC
AGTGATGATGAGCTTTTTTTCATGTTTGTGGCCACATAAATGTCTTCTT
CTGAGATGTGTCTGTTTCATATCTTTTGGCCACTTTTTGATGGGTTTTTTT
TTCTTGCAAATTTGTTTAAATTCCTTGTAGATTCTGGATATTAGCCCTTT
GTCAGATGGATAGATTGAAAAATTTTCTCCTATTCTGTAGGTTGCCTGT
TCACTCTGACAATAGTTTCTTTGCTGTGCAGAAGCTTTTCAGTTTAAAT
AGATCCCATTTGTCAATTGGCTTTTGTGCAATTGCTTTTGGTGTTCTAA
TCATGAAGTCTTTGCTCATGCCTATGTCCTGAATGGTATTGCCTAGGTTT
TCTTCTATGGTTTTTATGGTTTTAGGTCTTATGTTTAAATCCTTCTTTT
TTTTTTTTTTTTTTTTGAGATGGAGTCTTAGTCTGTTGCCAGGCTGGA
GAGCGAGTGGCGTGTCTNTAGGACGC

>Contig19

GCATGTTGTCTAAAGGTTTGTCTTCTCCAAAATTCATATGTTAAACCT
AGCCCCAAATGTGATAATTTTGGAGGAAGGCTCTTTGGGAGGCAGAGCC
CTCATGAATGGGATTAGTAGCCTTATAAAAGAGACCCCTGAGGGCTCCCT
TGTCCCTCCACCGTGTAAAGGATGCAACAAGAAAGTATGGTCTATGATCC
AAAAGCAGACCCTTGCCAGGTACCCAATATGCTGGCACTTGAACCTCCC
AGCCTCCAGAACTGTGAGAAATAAATTTCTATTTTTTATAAGCCACCGAG
TCTATGGTATTTTGTATAGGAGCACAACAGACTGATGTGCCACCCAAC
CATGATTATACGTGTAATTTATGGTTTCTCTGCTAGTAGGGATGCACCAT
GGGGTTAGGAACACGCTTTTCTTATTTCCACACAGTCTTAGCTCTAA
GCATGTTCTGTAATCAAAGATCCCCATCTTTTATGAATGAAGGAGTCAGT
GAATGAATTAATGAAAGAACTGATAACCCCTCAATAATTAATCCAGCCTTT
TATACCTACTATTAA

>Contig20

ACGGTTCTCTAAAGACTTTCAAGAGCTGGATTTTATGCTTTAGGTGAAGG
TGATAAAGTAAAGTGCTTTCACTGTGGAGGGGGCTAACTGATTGGAAGC
CCAGCGAAGACCCTTGGGAACAACATGATAAATGGCATCCAGGGGTGTA
TATCTGTTAGAACAGAAGACACGAAAATATATAAACAATATTCATTTATC
CCATTCACTTGAGGAGTGTCTGGTAAGAACTGCTGAAAAAACGCCATCAC
TAACTAGAAAAATTGATACCATCTTCCATAATCCTATGGTACAAGAAGCT
ATATGAATGGGGTTCAGTTTCAAAGACATTAAGAAAATAATGGAGGAAAA
AATTCAGACATCTGGGAGCAACTGTAAATCACTTGAGGTTCTGATTGCAG
ATCCAGTGAAGGCTCAGAAAGACAGTACACAAGACGAATCAAGTCAGACT
TCATTGCAGAAAGAGATTAGTACTGAAGAGCAGCTAAGACACCTGCAAGA
GGAGAAGCTTTGCAAAATCTGTATGGATAGAAATATTGCTGTCTGTTTTTA
TTCCTTGTGGACATCCAGTCACTCGTAAACAATGTGCTGAAGTGGTTGAC
AAATGTCTCAAGTGGTACGCAGTCATTACTTTCAAGCAAAAAATTTTAT
GTCTTAATCTAACGCTATAGTAGGCATATTATGTTTCGTATTATCCTGATT
GAATGTGTGATGTGAAGTGAATTAAGTAATCAGGATTGAATTCATTAG
CATTTGGTACCAAGTAGGAAAAAAATGTAAAGCCAGTGCTTAGACACA
GC

>Contig21

CGCTGTCTTAAGAACTGGGCTAGGAGTGAGCAGTGAGCCAAGATCGCACC

FIG. 4 (5 of 61)

59/118

ATTGCACTTCAGCCTGGGCAACAAGAGCAAACTCCATCTCAAAAAATA
CATATATATATATGACCCATAAAAAGGAGATAAATCAACACTTCAGAACT
GACCCAACTTGCAAAGATACTATAATTAACAGAAAAGGACAGTTTACTA
AGTACTCCGTATGTTCAACAAGTGAAAGATTAAACATATTAAGTAGAGAT
GTAGAAGATATAAGAAGATCCAAAATGAACTTTATAGAGTTGAAAACATA
ATATTTAAGATAAAAAATACACTAGGTGGGATTAAAAGTAGATTACACATT
GCTAAGATAAAAAAATGAGCCTGAATACAGCACAGTATAAACTATCT
TAAACAAAACACAGAGAGAAAAATAACTTTAGAGACTTAGCTCTTATC
CTCTATTTGTTTCTAAACAGAGGATAAGGGGCAGAAAAATGTTTGAAGA
AATCATGATTTTTAAATTTCCAAGTGAATAGGAATAGCACTGGGTAGTC
ACAGGAGGCTGGAAAGACCCAAACAGCAGTTAAACAGGAACTAGGCAAA
GAAACCAAAGGATAACAGTAAACCTAACTAAGGGAGAGAAAACTGACAA
AAGCTGACTTAGGATAACTGAC

>Contig22

CCTGAATATAAGCCGCAAGTAACCAATTAAATTTGTTTTCCAAAATTGTA
TTAACAATCTATGAAATTTTTATCTTGACCATAGCTATAACTCCAGAAG
CCTTTTATAACCTCTATAACCTTTATTAAGGAGTAGGTTAATGCTTCAAG
AAAACCTTGTTAATCTGACACAGGACCCATATGCTGATCTTGCATCAGTG
TGGCTTGGACATCAATGATTATGATTAATTTATAGAGAAATTGAACCTTAT
TTTATCTCTCAAAATTGGCCCTTACAATCTCACACACCCACCTCTCCAC
TATAGTTCTGGGCCCTTGAGTTGAATAGCTTTAATTTCTGGCTCTGTGT
TCAAGAATGCAGTTTATTTTGATTGGCATTCTTACCAGTCTGAAGATG
AACCTTTAATGCTGTGTCAGTATTTAAGATTTAGCAGGACTTGTCTTTTA
AGAACCAGGAGTCAAGCCCTATAACTCAATGTCACAAGGACTTTAAAAGC
ACATACATAAAGATATATGGATGTAATAATCATAATTTTTAAAAAATTGT
ATTAATCTCAGTGTCTTCTAAGCAAAACCAAACTTAATAATAATGGCATA
GAAATTATTTCAATAAAACATAAAATCTGTTAAGCCAGTTACCAAAAGGC
AAAAGAAAAGACCTTCTGCAATGCACAGAATATTATGTTGGAAGAAAACA
TTTCTTTAGACCTTTAAGAAAACATTGTTAGCATCAGGACACAACAAAC
AGAATCTGAGGGTAAAAAACGTATATGAGCTGAAGGGAGTTGAAGGAGGG
CATTACTATTTCCCAACCTTTTAAAGGGGAGAGAAAACCTAAAACAGCAA
GATGCAATAAAAGCTGAACCTTTGGGTTAAAAAAAATTCTTAAGTCTCTT
ATAATTTATTAAGAGTGAATCAACCCCGTAAGAAAATTTCAATTGTTCTAA
CCAAATTTTTAATATATAAGTAGTTTTTTAATCATCAACCCAATCTCTAGA
AAGACCATTATAATTTCCCTTTAATTATAGACAACCTTTATCATATAAAG
TTTTTTAAATAAATCCTCTTATTGTGACTTACACAGACTATTATGACA
TGCTTGGACTTTCTGGTTTGTGTCGTGAACATCCTTTCTTTCTTTCTT
TTTTTAAATTTTACTTTACGTTCTGGGATACATGTGAAGAATGAGGTT
TTATTACGTAGGTGTACATGTGCCATGGTGGTTGCTGCACCCATTAAAC
CGTCATCTATATTAGGTATTTTCTTAATGTTATCCCTCCCTTGCCCCC
CACCTCCTGACAGGCCCTGGTGTGGGACATCCCTCCCTGTGTCCATGTG
TTCTCAATGTTCACTCCCACTTATGATTGAGAACTGCAGTGTGTTTT
CTGTTC

>Contig23

GCTAAATATAAGCTATGATAAAACAGTTGGCCCTCTGTATCATGGGTTTC
ACAACCTGTGGATTCAACTAACTGTGGATGAAAAATCTTGGGAAAAAAG
AATGGCTGCATCTGTACTGCACAAGTGCGTGCTTTTATTCTCGTCATTAT
TCCCTAAGCAATACAATATAACAATTTTATATAGCATTACGCTGTAT
TAGGTATTATAAGTAATCTAGAGATGATTTGAAGTATACAGGAGGATGTG
CTTAGGTTACATGCAATATTATGCCACTTTATATAAGGCCCTTGAGCCT
CCTCAGATTTTGGTATCCATGGCAGTCTGGAGTCAATTCTCCTGCAACA
TCTCCATTTGTTTCAAGATTCTTCTATATCATGTTTATATCAGAAAATCT
ACATAAGATTTTTTAAATGTGTTTATATAGGTTTTGTGTTATTTTGGTGT
TAATCCCTAGATATATGCAGTATTTATTGCTATTATGAGTAGTGTCTT
TACCATGTATTCTAGTTGGTTATTGCTGACAGAGAAATGTTGCTGGTGT
TCTAAGTTACCTTGTCTTAAACACCTTGCTGAACTCTTATTAGTCTCA
TAGTTTTTAAATTAATCTTTCTTAGTTCTGATAACATAATCTGCAATAAT
GACAATTTTATATCTTTCTTCAATGCTTATATCTCTCAGTCCCTCTTA
TCCCAAAGTATTTCCAGGATCTCCACTATAACATTAAATAGTAATAAGA
ATTTCTGTCTTGTACTGATCTTAAGGAGAATAAATTTAAATTTCTCTG

FIG. 4 (6 of 61)

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TCAGGTTTTATGCTTGATATAGATTTGTGATATATAGCCTTTCACAGGT
AAAAAAAATGCTTTCCTAGTAGTCCTAATTTTTTAAAAAATCATCATA
AATAGATGTTGAACATTATCAAATGCTTTTTCTGCATCTATAGAGATAAT
CATATGGTTTTTTACTATTTATTAATGTAATGAATTAGACCAATTTTCTA
ATGCCAACTCTTTCTTGATTTGTAGGGTAAATCCTATGGGATCATAAAA
TACTTTTAATACATTGTTAGATTTGAAGAGTTAACGCCTTATTTAGAACG
TTTTTCAGTCACATCCATAAGTGAAATGGCACTATAGTGTCTATTACTATT
ATATTTTTCTGGTTCTGAAACCAAATTATACTCACCTCATACAGTAAGT
TGGGCAACTTTTGTTCTTTTTCTGAAACAATTTGTGTATAGAAGAAAT
TAACTGTTCCCTGAAAGTTTGATAATAATCATCCAGAAAATTATCCCAT
CTAGGGCTTTTACAAAAGGAGACTCTAGAATGCCATTTTCGGTTTCCTTG
ATGTGTATTGGCCTCTTTCAATTAGGCTTTTGGATTTTATAGGGCATTTT
TTCATATAGGCTTTTTACCGG

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CATAAACTTCAGGTTGGATGTTTCGGTCAAAGTGGTCCGGCGATGCGAAAA
CGAGAGGGCTCGAGGACTGGGAGAGAACTATTTGAAGGTATCTCTCAGG
GGAAACCAAGCGGAAGGCGGGGAGTAAATTGGGAGGGAGCGAGGCCTT
CAAAGAAGGGGCTTGCAATTAGATCGGCGAGATCCGGGAGGGTCTGGTGGG
GAGAAATGACTAGAGGACAAATCTAATGGAGAGACAGACGGAGATAGATA
TCGTGACAGAGAGAGGGACAGTGACAGCGCACAACAGTGCAGGGTCCATG
AGTACAAGGCCCTTAAGTGTACACCCAGCCGGAGTCATGGCAATTCGAT
TCCTGTACTGACCACCCAGGATTTGGGTAGACTGTACGAGTTAATGAGCA
TGGTCCCCAACAAGACTGCTTCGACCTCAGATGCAAAGCACACTTCAGGG
GTCCCCAAGCCACTCATGTTTTTGAATGACTGCCATAAGTTCAAAAATT
CCCACAATTCTCTCAGATTCAATAACTGGGTATAACCACTCATAGAATC
AAGAAAATGCTATCATTATTATTACAATTTTATTATAAAGGATACAAATC
AGAAGGACTAGCCAAATGAGGAGACACATAGAGAGAGGACTAGTAAAAAA
CAGAGCTTCTGCGTCTACCTTCAAGGAATCAGGATGCACCACCTCCCA
GCACATCAAGTGCTCATCAACCAGGAAGTTCTCTGAGCTCCAATGTCCA
GAGATTTTAGGGAGGATTCAATACATAGGTATCATTGATTAAATCATTGG
CCATGTACTTGAATCAATCTCCAGTGTCCCTCTTCTCCCTAGAGGTCTG
AAGGGTTGGCTAATCATGTGGCTCAAAGCCCCAACTCTAATTACCTTT
TTGGTCTTTTCAGGGAAGTACCCCATCCTGAAGCTATCTACAGGCCCTG
CCATGAGTTAGCTCATTAAACATAACAAAGACACTTATATTACTCAGAAAA
TTCCAACAGTTTTTGAAGCTCCATGTGAGGAACCTGGGACATAGATCAAA
TTCTTTTTTTTTTTTTTTTTTGGAGACAGGGTCTTGCTGTGTTGCCAG
GCTAGAGTGCAACGACAGATCAAGCTCAATGCAGCTTCAACTTCCCAGG
CTTAAGTGACCTTCCACCTTAACCTTCCAAGTATCTGGGACCAAGAAA
ATGGCTAATTATCCTGGCTGATTTTTAACTTTTTTTTTTTGTAGGGATG
GGATCGCCCTGTGTTGCCAAGGTTGGTCTCAAACCTCTGGGTTCAAGCAA
TCATTCTGCCCTGGCCTCTGTGATGGTTAATACTGAGTGTCAACTTGATT
GGATTGAAGGATACAAAATAATTTTTTGGGTGTGTCTGTGAAGGTTTCG
CCAAAAGACATTACTTTGAGTCAGTGGACGGGAAATCCCCCTTCCCA
TGGGACGGGAGACCCCCCTCCATCCAGGTAAAAAAATCTAATCACCTGC
AATGTGGCAGAAATAAAGGAGGGAAAAAACGGGACCCCTANATGGGTTA
TTCTCCACCTAATCTTCCCCCAGG

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CCATGTATTTTCAATTTCTACAGACCCTGAGATGAATTTGTCAATTGCCACGG
GGTCTGAAGTTCAAATACTCTATTTGGTATCCTGCCCTGTGGTTAACT
GTGATCATTTCACTCACCTTGTTTATGATGAGAGGTGCCACCATCTGGCC
TCCTCCACTCTGCAATCCTGTTAATTCCTATCAAAGCTGAAAACCTGCTG
CAGCACCCACACCATCACCTCCAGCCTAGAGAGGGGAAGCTACCAGTGAGC
TCTCCTGGATGCCGGTGTGCCCTCGCCAATACATTTCTTCTTAGTCCCT
TGGTCATCCTGAGGTGTGTGATTAAATGGACAGCTATGTGGATTGCACATA
ATAGATGTATCCAGCATCTTCATCCCTGATTTTCTTTTACAGAAATCAC
TCAACCTTAGCAACATGTGAAAATCACCTAAGGACATTCTTTAAATCCCT
CTGTCCACATGGCAACACAAACCACTTAAATAAGAAATCTCCAGGGAGTCA
CTCAAGCATCAATGTTTTTAAAGCTCCAATTTTAAGGATCATTACATTA
TGTCGAAGAAATTATAGTATTTTACGCTTACTGACTGTAAACCACCACCA
TATCTAAGCATCCATTAGTCAACCTAGCAGACAATAAACTAACATTACCT

CCAGGTACTCAAATCAATTCATTGTCATCCCAAATCCCAGATGGGCCACC
CTTATTGACAAATTCAGCCCAATCTTGGTTGAACACATTTAGAATATATT
TCCATGAACAATATCCGGTTGACGAGTTTCTTTAACTTTTGGAGTTTAA
GCCATTTCCCTTTCACAGTAGCCTTGTTAATTCCCTGTCAATGCTCCATGG
GGGTCAATGAAGAGACCTCTTATTAAGTGTGAAGCAACTTGGCTCAGGTGC
AGACACTCAAATGCTTCACATGCAGTGGGAAAAGAGAGTGATTGTCTAC

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TTTAAAAAGAACTGAGTCTTTATTTCAGTCGATTCTTCTAATCTATGAACA
TAGCATCTCTCTCAAAGCATTTAGTCCTTCTTTAATTTCTGTCAATTAATT
TTTTAAATTTTTCATCCTAAAGATTCTGTATATGTTTTGTTGAATTTATG
CTTAAGCATTTCACCTTCTTGGTAACAATTATAAATGATTTTGTGTTTTT
TATTCCACTAGTTTCATTTTCAGTGTGTAGAAAAGCAATGAATTTTGTGT
GTTGATCTTTGTTCCAACATCTTGAACATTATTGAACTCATTATTAGT
TCTAGGAGGTTTTTTCATTTTTCTTGTAGATACCTTGAGATTTTCTATAT
AGACAGTCATGTTGTCTGCAAACAGGCACAGTTTTATTCTTCTTTCA
ATCTATATGCTTTTCTTTTTTTTTTGGCCTTATTGCAGTGGGTAGAACTT
CTAGCACTATGTCAAATAGCATTGGTGAAAGCAGACATCCTTGTTCTTGT
TCTTAGAGGAACATTTGGTCTTTAATCTTGATTTAAAAAATTCCTTGCAC
TAAGTTACCGTGTGTTTGGGGAGGGAGAGGTGGGGTGAGGTGGGGATTTC
CCCTAATGTTTACAAGCTGGGATTTTCTTTTCTGTGTCTAATTATTTT
CCTCATTTGGCTTGAAAAATCTGATAAAACATTTTAGGACTGTGTATAAAA
TAGAATTAGCCAAGTGCAATGTCTTTATTTCAGAAGAAATTTTCATGGACGT
TGTGCCTACTCTCTTGGCTTCTTGGCTTCATGGCTTTCAGATCCACAG
TAAGCTCTGGATAGTAGAAGTTATAGTAAGACTGACTTCTAAATAAATGA
AGTGACTTTAACCTTACTGATATGGCTTAAAGAAAAGGAGTGGCCTTTAA
GATCCATGAACCTTCTCAAACAAAAGTGATAACGTTATCTCCATGCATATA
TAATACTAAATATAATGCAACTGAGAGAAGTAGGCTGTGGTAAGAAAGGA
GACCCAAGTGCCATCTGAAGGCAGCACTTACCCTCTGCTTCATCCCACC
GAGGAAACAAAGCATGAGTATTGCCAGATTTTCTTCTGTTTCAAGAAAAG
CCAGAAATCCAGGTTTTTGGCGTAAATGTCTGATTTTAAATGTTGGGAAC
TAATTTATATTTTGAATAACATTGTGTGGGACAAGTGAACCTGTATGTG
GAACGTGCTTTCTCCAGTGGCGACCAGTTTGGACCGTTGATACTCAGCAA
GTTTCAGCCAAGTGCGCCTTGTCTATTGTCTAGTCATCAAGGTGATGTGTAT
TGGTCAAGCAATTAATTTTGTCTCAGCATCTCGTGTGTTTTCAAAGAAGT
GAAGGTTTCATTTGC

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TTTCAGAGCACAATGCGTATTTCATAGTATATTGACTTAATTTCTAAGTGT
AAGTGAATTAATCATCTGAATTTTATTCTTTCAGATAGGCTTAACAAATA
GAACATTCGTGTATATAAATGTGTAAATTAGAGTTAATCTTCCAATCACA
TAATTCGTTTTATGTGAAAAGGAATGAACGTGTTCCATGCTGGTGGAAAG
ATAGAGATTATTTTATAGAGTTTGTCTGTTGTGTTTGGGATTCTGTTTTT
TTTTAAATTTGTAATATGTACTTGTGTGAATGATTTTTTAAATGATTT
TACCATTTTTGGAAGGGTATTTAATGATAGAATATCATCGAGCCAACATG
CACTGACATAGAAAGATGTCAAAGATATATTAAGTGTAAATGCAAGAGG
GAAAACACTATGTACAGTCTGAGCCAAATCAAAGCATGTATGTTTTTAT
ATGTGTACAACAAAAGGTTTGGAAAGATATGCGCCGAATTGTTAAATGTG
GTTTCACCTTGAGGGGGTGGGAGGATGGGGCCCCAGAGGGGTTTTATGGG
GGCCTTTCACTTGGTATTTTTTTCATTTTGTCTGTTTGAATTTTGTGTT
TTTCTTTTAAATGGAGTTTCACTCTTGTGCGCTAGGCTGCAATGTAGTG
GCGTGAACCTCAGCTCACTGCAACCTCCGCCTCCAGGTTCAAGTGATTCT
CCTGCTCAGCCTCCCATGCCTCCTGTGTAGCTGGGATTACAGGCACCCA
TCACCATGCCTGGCTAATTTTGTATTTTTCAGTAGAGATGGGGTTTCACC
ATGTTGGCCAGGCTGGTCTGTAAATTCCTGACCTCAAGTGATCCACCCACC
TTGGCCTCCCAAAGTGCTGGGATTTTCAAGGTGTGAGCCACCACGCCCAGCC
CTGTTTAAATTTTATAAGTATGTACTACTTTTGTAAATCAGAATTATTA
GAAAGCATTTTACTGATTTTAAAGCTTAGACATGTTCAAATGCCTGCAA
ACTACTTAACACTCAGCTTTAGTTTTTCTAATCCAAAAGGCCGGGCAGT
TAATCTTTTTTGGTGCCAATGTGAAATTTAAACGGTTTTATGTTTTTCTG
TGTGTGAATGAAAATATTTCTGAGTGGTGGTTTTTTCAGAGGTAGACC
ATGCTCTGTCTTGTGTTTTCAAATAAGTATTTCTGATTTTGTAAATGAAAT

FIG. 4 (8 of 61)

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ATACAATATGTCACAGATCTTCCAATTAAGTAGTAAGGGTTTATCCTTAA
TCCTTGCTAATTTAAGCTTGCATAAGTCACTTTACTAAAAGATCTTTGTT
AAGCTAGTATTTTAAACATCTGTCAGCTTATGTAGGTAAAAGTAGAAGCA
TGTTTGTACACTGTTGTAGTTATAGTGACAGCTTCCATGTTGAGGTTCT
CATATCACCTTGTATCTTGAAGTTTCATGTGAGTTTTTACCATTAGGATG
ATTAAGATGTATATAGGACAAAATATTAAGTCTTTCTTTTACCTAAGTTT
GCTTTCTTGACTAGTAATAGTAGATATTTCTGTAATAAATGTTCTCT
CAAGATCCTTAAAATCTCTTGGAATTTATAAAATTATTGGAAAGACAAGA
ACAGTTTTTTATTATTATATGCATTATTATCG

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CTTTCTCAAGAAAAGGGAAGTGGAGCAATTAACATATGTAATTTTTTTT
TAAAAACCCCTAAACCTAAACATCTACCTATATACAAAATTAATTAACA
ATGGATCATGGACTCCAATGTAAAACATGAAACTCTAAACTTCTAGAAAA
AAAACCTGGAGAAAACCTTTGGTACCTATGACAAGGCACAGTTTTTAGACT
TAACACTAGAAGTGTGAACATAACAAGAAAAATTAATAATTTGAACCTT
ATGAAATCAAATTATTTGCTCTCCAAAAGACCTGTTAAGAGGATGAAA
ACTAAATTACAGATTGAGAGAAAAATTTTGTAAATCACATATTTGACAAT
GGACTTGTATCTAAAATATCTAAAGAACTCTCAAACTCAACATTAAAAA
AAATATCTAATTAGAAAATGAGTGAACATTTTACGAAAGGGCCTTATAG
ATTAGCAAATAAAACACTTGAAAAGATACTCAGCATCACTAGCCATTAGA
AAAATGCATATTAACCACAATAATGTATCGCTACACACATATAAGAAT
GGTTTATGAAAAAATAGTGATGACCAACTGTTAGTGAAGATGTGGAGA
AACACTCATACATTGCTGGTAGAAATGTAAAATGGCATAGCCACTGTGGA
AAATTATTTGGCAGTTCTTTTAAAACCTAAAAATCAATCTACCACACAAC
CCAGCAATTTTATTACAGGGCATATATCCCAGAGAAATGAAGATTTATGA
TCACACAAAATCTGTACACAAATGTTTTATGGTCACTTTATTCTAATA
GCCAAAACCTGGAACTATCCAAATGTCTTCAATGGGCAAAGGATTAAA
CACACTGTGATACATCCATACCATGGAATACTACTCAGCAATAATAAGGA
AAGAATTACTGCTACACACAAGTTGGATTAACTCAAGGAAATTTGTGCTG
AGTGAAAAATTAACAAGCCAATCTCAAAGGACACATACTTCATGATTCCA
TTTGTATAACATTAATTAACACAAATTAATTACAGAGATGGAGAACAGAAT
AGTGGTTGCCAGGGATTATACATGGTGGACGCGGTGAGGCGGGCCTCCAC
GCCTTGGAGATGAAGGGGGCTACACCTTTTAAAGCACACCCACGAGAGAG
TTTTGTGCGGAGGGGGCCCAATTTAAGTACTCCGCCCCGGGGGGGAACAC
AGGGGCAACAAAAAAATTTGGCCTTGGGGGTGACCAACACACAAAAA
AAAACAAACACACAAAAAAACAACNATGGGTGGGAGGATTAATCGCCAAA
TCTGAGTAAGCTATCTGGACAGTACCAATATCGATTTCCAGTTTTGATG
TTGTACTATAATAATGAAGATGTTAACAATTGGAAGAAGCTGGCTGAAGG
GGGCTCAGGAACTCTCTGGACATTTCTTTGTACCTTCTGTGAATCCATC
ATTATTACAAAATAGGACATTTTCTAAAGGTTAAATCATTTTAATTTTAA
AATGTCCCTGTTACTGTTGAACTCACATCTCCATATACTGATCAAGAAC
AGCACTAATGGCCCCCTGGCCTCCAGGAATTCACAAATCTACTGACTTTT
CTTTGAAACCTTGGCCAAGTCGCTTCTCTCTCTGCTCCTCAATTTTTCA
TCTTCAAAATGAAGATTGAATGACTATTAATCTCTTGCAATTTCTTGAG
ATGAAGGGTCTTAAAGGAACTGAAGAGGATGCCATGTAATGTAAATATGG
GTTTTTACTCCATCAGCCAGCCAGACAGAGGGCAGACCAAGACATGG
TAACCAAGGAGGCCATGTGTAAACAAAGACCATTTAGACTTATGCTCTGG
CCTTTGCAGCCCAACTGGTGTGGCCAGTTGGTGGGGTATGAAGAAAATGG
GGCCTTCCAGGAACCATGTTGAGTGGAGATAAGCAGGGAGGAATGCAGAA
GACATGGGGGCAGTGCCAGTCTCAGCCCCAGCCAGCTACACCCACACATG
GTTATGAAAGACTGACAGCCTGTAAGNTGAACACAGCCCTGCCTCTCTTA
GATAGGC

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GCAAAATATGATCTCAGATGTGGATTTACTGTAAAGTTCATCAAATTTAAA
TTTCAGAACACTTAATCTGCAAGAGTCCTTTCCAAGACCCTATACCTAAT
TTTGTGTTTACAATTTTATATTTGTTTTCTTAAAGAAGACCACCAATATA
AACTATATCCAGCCTTCATGATAAGTACATAGGAACTATGCAAAATAGG
GGGAAAAAAAACAAAGAAAAATACCTAGTTTACTAATGGTTCACCTCTGA
ATAGCACATATTATAATGATACAAGCACTCATTACTAGTCTAGGAAAAAT
GAAGATATAATTGCATTAGGAAGATCAAGAGGTAGGAAATGTGGATGTGT

FIG. 4 (9 of 61)

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GTGGTATAGACTAGGGCAGGACAAAGAACCTAAATCCTCATTCTTCTAAAG
ATAATTGTTAATACGTAAAACTCAAAATTCAAGAAGTAACAGTAAAAGCG
GTCATTAAAGAAACAAGCACTAAACACCAGATAGGAAGCGAGAGATGGGGG
AAGAGGGCGACAATCTGATTATTTTTTGCAACAAATTTTGTAACCATT
TGA CTGTTTACATGTAGAACTTGGATCTTTTTTAAAAAACACAAAATAAT
AATACTATTATTTTTTAACTGGATTTTTGAAAAAGAAGATAAAAGTCTCA
TTTTAGTAATTAAGCACTATTCCAGGTTAGTCCACTCAAACTTATATTC
GAAAATTAAGCACTTTGGGAGGCTGAGGCAGGCAGATCACCTGAGGTTGGG
AGTTTCGAGACCAGCCTGACCAACACGAGAAACCCCGTCTCTACTAAAAA
TACAAAATTAGCTGGGCGTTGTGCATGCCTGTAATCCAGCTACTCGGA
GGCTGAGGCAGGAGAAATTGCTTGAACCCGGGAGGCAGAGGTTGCAGTGAG
CCGAGATCACACCATTTGCACTCCAGCCTGGGCAACAAGAGTGAACTCCA
TCTCAAAAAAAAAAAAAAAAAAAAAATTAACCTCTGGAAGTTGAGTTTG
CAAAATTCATTATGCTCATTTTTTTAACTTGATGTTTGGAATGTCTG
ATGAAAAATTGAGTTGGGGGATGAGAAAAAGAAAAACATCAACCCAC
AGCCCATTCATTTTTAGCCCGACCCACAGCTCCGGGGAAGGCGAGCAGG
TCCATCCTTCACTCTTTCTTACCTCTTTCCCTCCTTCTGGCTCTTCCA
CCTCTAATTTGGAGCCCAAAAAAGGCACTGGGAAATGGAAAGTCTTTT
GTACGTGGTACTTGCCGGGGAAGCTGCCATGAAACCTGGCCCCACGGTG
GGGAGGGAATGCCANCTGAGGCCTCGTGCCCATGCTAGGATAGACTCGT
CCAAACATGTCAAGTGGTCTGACAGGGCAAGCANCANGAAATCATGTATG
AGTATGAAGTATCTGTATGCAAGGGCGGGGAGAACACGCGGAGGAATGG
GGCGTGAGAAAAACAGCACAGTACGTTTTCTTAGCAGCTGTCTCTGCTCAG
CCATGGGAGGTACAGAGAAAGAGGCTTGGAGGCGTTATTTTCACTGTGA
GATGTGAGTGTAAAAAGTGCCCAAGACACAGTGAGTACCAGGGAGATGC
CCTCTTTCTACCCGAATGCAGAATGGCCACAGGCCTTAAACACACACA
TGGGTCTCTCAGAGGAGAGAGGCCTCCACAGTGGACACCCGCATTCTCCCC
TGGTCAGCAGCAGCAGGGCGAGTGCTGGGCCATCATGAAGCTTTCAGGGC
AATGAGCTCTCAGCAATAACAGGAACAGTGCTGGGGGACTGTAGCTGCA
AGACCGATTTTCATGTAAGATGGCCTCTGAGGACTCCGAGATACACCAGG
CTGAGACTAGCTGGCAGCTCCAAGTTGTTGGTCAGAAGAGAACAGGAAT
AGGGAAATTGGAATTACTGTTACTACAATTCCTTTACATCCGCACAACCA
TGAGGTCCAGCGATTTTCTATTATTTTTTTTTTTAAGACAGGGTCTCAGT
ATGTGCGCCAGCATAGAGTGCAATTGATGTGATCATGGTTCACTACAGTAT
TCAGTCCCAGGCTCAAGTGACCTCTGCTCAGCCTCTCAAGTGGCTG
GGACAGCAGTTGCATGCTACCAGGCCAGGCTTTTTTTTTTTTTTTTTTTA
GTTCTGTAGAGCACATAGC

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GGTTAACAATGGCACAGGGAAACAAACAGTTCAGGTGCAGGGGCTCTAA
ATCTATCATAAGATGTTAGGTATGGGGGCTCTGCCGGACACAACTCAAG
GCTTTATGCTGTTATCTCTTGAGCGAAATCCTGGGAACTTCGTACATTGC
TTGCTTCAGTACCTTATCAGTTAATCGGACTCTTTGATATGTTGGGAGTC
AGCGTACACAAGTTAACTCCTTGAGGAAGGGGGTGGGTAAGGAGTCCTTG
ATGTCTGGTAAATGAAGGAGCGAAATCGAGTTCCTCTGGCTTTCTCAGCT
AAGGGAGAGCTTATTATGTGGAACAAGGCTAAGTGATTAAGGGAGAAA
GGGAGAGTCTGAAAAACAAGGTTAGGTATTACAATGTCAATAAAATTGGTC
TCCTTATACAGTCTATGGTAGATTCTTTCCATCTTTAATCTCCCTCTA
GCACCACCAGACTTTTTCTCTGTACCTTGAGATGTAAATTTTGCTATC
TGAATTTTCGTCTAAGAGTTGTTTCCTTTAATATGCAATTTAGGGTTAT
TTAGCTGACAACTGCCAAAGTAGTGAAACAAGTTATCAAGAACTTGAACG
TCTAAGGTAGGAAAAAAAAAAGTCTTTATGAATCTATAAGATGTACTCT
ATTGGCATGCCTAATACGTCTATGTATTTACGTGTTGTGTACACAGTTTT
TCACTACTGAAATATATAGAGGAGTTCTAATTAATTGACTTAAGACAAT
AAAAGCGCTTGAATCAATACCTTATCAGGAAAAAGGAAAAGACAAGTCA
AATGCTTGTTCAGTTTATATAACTTAAGTAAATCTTTAATAAATAAGC
TAGCTTTAATATTATTTGAAATGTCTTAAGAATTGCCAGCAGGTTCTGGG
TTACAGAACTAGTGGGGGTGAGTGGGGTGAGGGTTGGTGGGGTGGGGG
TGGTACGGGGGCTTTGTTTTTTCTTGCTGCCCCCTTCTGGGTGGGGGAG
TGGCAGGACCTTGGCAGCACCCCGAGCCGGCATGGCGTTAATAATGGAGG
GATGCCAGACCCAAGTGGCTAAGGCCCGGCTGCAGAGCCAAGTTGGCATT

FIG. 4 (10 of 61)

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TCCAGACTGGGGCTCGGGCCGCACCCCTCTCCAGGACCCCTCCCCTTGTACC
GAGCAGATTGTCGCGGGCAGTTTGGGCCAGCTGTCTGGCGTGGAATTC
CCAAATTCAACAAATCCTCCAAGAAATCAATCCATCCATTCCATCCCA
TCCATCCATCCATCCATCCATCCATCCATCCATCCGTCGAGATTATGAAGCAT
GGATCATTACTTTTGGGATGTGGATATATTTCAGTTAACAAGGAGCAGCTT
TCAAGAGCTGGATTTTATGCTTTGGGTGAAGTTTAGAAACACTAGCTCCC
AC

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ACCTCATGTGCTCTAGCGCCTCTTACCTCATGCCCTCCACTCTCAGTCTT
GCACTCACCTGCCACACTCAAGGGCTTCCCAGGTTCTTCTTAGATTCT
CACCGATAGCTCAGGGACTTTGCACATGCTACGGTCTCTGCCTGGCTCCT
CCCCAGATCTTCTCATGCCTAGCTGCTTCTCATCAGCACCCCTCAGAGAC
TGTCCCTGCCCCACCTCTCCAGGTTCCATACCTGCCACCCCTCCCCAATC
ACGTAACAGTTTCTTCCAGAGCGAGTTACCATCCCAGTATTTCCCTAAC
TTATTTTTTGTGACTGGTCTGTTGCCTGTCTCCACCACAAGAACATAAGC
TGCATGTGAACAGGAGCCTTGTCTATCTTGTACCCCACTGCTGTGACA
TAACCTGATACACATTAGATGCTCAATGATGTTTGATGAATGAAGTGCTG
GTAGTCCAACCTGTGTTTCTTGTCTGTGTAAGTATGTCTGTTGTGGTTTC
CTAAGAACCCTACAGCTCTCCCACTGTGACTCCTGTTCTATGGTCTGATT
TGCTGGACTAGAATCCTAACCTACATGCTTACTCTTAGTGTCTCTCCCA
GAGGCTGAATCCAGTCCCTAACCTCCACCAATGGCTAAGACCTAGCT
TCCAACCAGACAGGCCTACGCTGAGACCTCAGCACCGCCCTTCTGCGGTC
TCATCCTTAACGCATCCTTCAGGGCCAGCTTAAATGTCTCTTCTCCAAG
GAAGGCTATCCTCTTTCTGCCCCCTCAGTGCTCTCCATGCCCTCTCTATGC
CTCCATGCCCTGCTTTCCAACCTGCAGAGGTGGAGAAGTTGCTAATCTGC
TGTGTTGACATGTGCTGGGGTGCCTTGGGCCAGGGAGCAGGCTGGTGGTG
TGCTGATAGCCCCGTGGCTGTGCCAGGTCCATGCTCACTTCTGAGCCCC
AGTGGAGTAGGCTCCCTTTCCCTTATTGCAGCACTCAGAGGAAGGACGTG
CTTCTTAGGACAGATCTGGCCAACCTCTCCCTCGTGAGAGAAGGCCAGC
CATCCTCTTGGCCTCTTTCTTCTCTCTGCCCCGAGTAATAAAGGTGCCT
GGTCAGAGCCTTCTAGAAGGAGACCCAAACATCCACCACACATTCCCAGT
TCCAACCGTCATCCACATGGCTGGCTGTGCAGGTAAACGCAGAGTCTGTT
TCACACACCCAACCATCTAGTATTGGATGGGAGGACAGTAGCGTGACACT
CTTCTCAGCCTTGAGCCCTACTGTGGGCCCCACCCAACCCAGATACCAG
AGGAGCCCTGTACTGGGATGCTATTGGATGCTTGTCCAGTCATGTACAAA
GTTAGCCCTTGTATATAGAGTTAGCTACGTACATCTTCTCTGTAGGG
AACCCAAGAGGGGAGAAGAGATATGTAGTAGGATTTAACCTGCAATCCT
CTGCTGAGCACCGTGCACTACATACAGTGGGTAGCATGTGGTAGGTGCTC
AATAACTATTGACCGATCTATTGAATACAGTAAGATCGTGACACTATCT
AAAACGNGGGGTGTGGGGGAAAAACCCCCCTTGTTTAGGAAACCCAAA
TTGGACCGTGTGGC

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GCGCGATTGTGCTAAAGATCATGCATGCCTGATCAAACGTCCCCATATGG
CGTCTCAGAGTCAACTCCTTCCCCATCAGTGCCCTGACTTCGGCATAACA
AACCTGGCAGGTTAAGTGATTAATCGGTCTGTACAACTGTAGCCCTTAG
CAGGAAGCACTAAGCTTTCGTTTTCAATTTATTTCTTCCCTGGAAGTCAAG
AAATGAGGGATGCCTTCCGCCATGAAGTTTGTCTGATGTCCACTTTGTT
CTCAAGGAGATATTACAGTTTTTAATTTGTCTTTCTCTCCTGCATGGTC
TCCAAACCTGTCCAAAGAAGCCAGCTGGCTCCATCATCTGTAAATCACC
ATTGTCAACAGAGCACTTGACTTCTGTGCCCCATAATCCACCTGCACT
TTATTTCTGCCACCATGATAATGTAGTGTTACTACATTTTACATTCAGC
TGTAAGAAATGTTACATTCATTTACTTAAATCAAATTAAGTCTGCTCACT
CAGTCCCCCAGTGACCAACTTATAAAGAGAAGGTACATTTCACTCAT
CACTGAGGTTCTCTTACCACTGGAAAACCTGAGGAAGGGTCTGGAGTCCA
CAGTGGTTAACATCATTGCCTCTGTTTTTCTCTACTCAATGTAACCAT
CCAAGGTTACTCACAAATTCACAAAAGAGGTCTTCACCTCTGCTCTCAA
GACCCAGAGGGCTGGGTTCTAACTCAAAGGCAATGTTCCCCAAGTTTT
TGCATTGTTTTCAACATTGGGGAAAACCTGAGGGGATTCAAGAATGGTTAT
ATAAGTTTTGTGAAAATGTATAATTTTTTAAATTAATAACAAAGTA
TTATGGAAAGCACTAAATATTGAATTTATATAAATATTCCAAATATTTTT

CTAAATTTTGTAGTGAGAACTTGAGCTTGCTTCTGTGAGATATTTATTTT
AAAACAGATTTTGACACTTAAAAATGTCTAATCAAGCCTTTTAAACCATGAT
CTATCTCTTCAAATTTCTTCAGATGCCACCATCAATAAAGAAAACCTTTGTTC
ACACAAGTAAGTGGTAGCAAATGGCAGGGTGTATTATCATTTTTTTTTTTT
CTTTTTTTTGAGACGGAGTCTCGCTCTGTGCGCCAGGCTGGAGTGCAGTGG
CGCGATCTCAGCTCACTGCAAGTTCACCTGCTGGGTTCACGCCCTTCTC
CTGCTCAGCCTCCCGAGTAGCTGGGACTCAGGCACTGCCACACGCC
CGGCTAATTTTTGTATTTTTTAGTAGAGACGGGTTTACCGCTGTTAGCC
AGGATGGTCTCGATCTCTGACCTCGTGATCGCGCCCGCTCGGCCTCCCA
AAGTGCTGGGATGACAGGCGTGAGCCACCGCGCCCCCGCGTGTATTATCA
TTTTTGCCTGATGAAATTTTCTTGCCACTACTCTGGATGGTTTGATAC
ATTTAAATTGTGCTTCCAGGGTACAATTATCCTTTAAATCTATACCTCTT
TCCTTTCTTTTATTGACAAATATAATGTTACACTTTTCTGTCAATGACAGC
CACACCACCAGTACACAGATCCCAACAGAGTTGTAATATTTTATTAGTTT
CAGAGTTTCAATATTTTATCACTTTCAATACTTCATGTGCAGGAGTTTTA
TTTGGTACTTCTTTACAAAATAAATAGTAGTGCTTCCAAGCAATTTCTTTT
AATAATTTCCAATCAATGTTATTAACTAGTAATACTAGTATCTGTTTATT
CATAAATTCACAGGAAATGCTTTTTTACTTATTAGTCTTTGGAATTCTGT
TGTTTGATATAAACATCTTTCATGATGGCTTTGTGTCTACCAATAGCACTA
TTGCCAAAAGGCACCTTTTTCTGTTCCTTTACTTCACTGGTCCGAAGCC
TGGTACCAACAACTACCACACAGACTGGGAAATGAGCAATTTTGCCACGT
GCCCTTAGCTATTAATGGTGGCACTCCATAACTAGCATCTTAAGTCAAT
TTCATGAAAGAAATGTGTTTCTTATTTTGTACTTGCAAGGCACTTTTTAA
CTTGTAATCTTTTATTCTATACTTTTAAATTTAAACAGAGTAATAGAACC
ATAGAAGGAAATCAATACCCACGAGTCCATACTGATATAAAATAAATAGTT
ACATAAATAAATGGGGGGAGAAATAACAGCTCTTCTTACAGAAAAATTT
CAATTAATAAATGAAGAAGGAATTAGGGAATACAACGTTACCATTAAGC
AACCACAGTAATAATCATTACAGGCAATATCCAAAAATAAATCCAAAGC
CAGTGGGCAAAAGTTTGAGGAGATACAGGATATTAACATAGTCTCAAAT
AGCTCATGCTATTTATAAAATACAAAAGGAAACATAACCACTGTATAGTG
AAGAAACTCAGCAGACACCACCTTAGCGAAGTGATCAAGGTTAACGTCAC
TAGTAATAGGCTTGTGTGACATACTGGACTCCAATCTGATACACTGATAA
GGACATGACTTCTGCAGTATTCTTACCAAAACAGAATTCTAATGTAA
TTAAGGAAAATGTCAGACAAACCTATTCTGAGAAACATTCTATAAAACAA
CTAACCAATACTTTCAAATTTGTCAAGGTCATAAAGACCAGGCGATGGTC
ACAGATTTGAGGAGACTAAGGAGATACAACAATAACAAATGGAA
CCATGGCATTCTTGATTGGATCTTGAACAGAAAAGGATATTAGGAAGA
AAAGCTGATGAAATTTAATAATCAATCTGTAGTTTAAATTAATAGTATTGTA
CCAATTATTAATTTCTAGATTGATCATTATACTATGGTTAAGTTTTTAA
CATTAGAGGAATCTGGGAGAATGGTATATATGAACTCCACTGTTCAATCA
ACTTTTTTCAGTAACTATTATTTCAAATAAAGTT

>Contig33
GGGAGCGGGCGGCCACGCTGATCTCTAAAGCTTTAGACCACATTGGCTCG
AGCATGGTCATGGCCGTTTCCTG

>Contig34
GACGTCCTTAGCGCTATATTATAAAGAAATATTCACCTCCCTGCTGAGCTT
ACAGGGTGTACCTAATGTCCAACAATATGAAATCTCTTCAATGAATTGCA
GCACGTCCATATATAACCCACATGGAAGCTGTCTCTTTCCTCACTTCG
AACTTCCCATGCCAAGAGGGACCTCTTGGACTCAAAATACATCTTAGCAA
TATAGAAGATGCTGGAGACTGTGAGGAGAAGTGGAGAGGGTTTACAGTGT
AGCCCCACAGAAAACAACTTATGACCCCATCAGTCACTTGTCCCTTTTTT
CCATGCCTCAGTCTAGTCAGGAAACCACTAGATCCTGGATGGCTTCTTCT
CCCTTCCCTCCTTCTCTTCTCCTCTCCCTCCCTTGCTCCTCCTTCCTC
CATCACCCACTCCTTACTTCCAACCAAACTTGACTAGCTCCAGTCTCAT
CCCTCCTTATTGAAAACTATTTTACTCAGCCCTCCTCCCCCACTCCTGCC
CAATCTTTATTCTTACCTACATCAGACTTACCAAAAACAGGCCAGGA
TAATAAACAGGACAAACTTTTCAAACACATTTTAAATGACCATATTTTGT
TATTTTGGTACAATTTGAGGAGTCCCAATCCCAGGGAAGACTAACAAGA
AGTTCTCCTAACAAAGGTGGGTCTCCCTTACTAAAACTCCTGTAATGG
CTGAAAAGAGCATGAGGTTTTCTGCATATCATTACATTCAATAGAACG

FIG. 4 (12 of 61)

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TCATGCAGCTGTTAAAAATGATCTGTAG. AGGCTATCTTGTGACAGAAAG
 GCATTGGAGATATACTGTTAGTGACAAAAATAGGTTATAAATGAATTTTT
 CCATGCATGCCTCTATATTTATAAATACACACACATAAAAGACAGGAAGG
 ACAGACATTAAACATTATAGTGCTTAAGATGATGCATAGTATAATAGTT
 AGGACCATGGCCTTTGGGACAGAAACTACAGCCTCTCTCCCACTTATCA
 GCCATGGGACCTTGGGCAATTTGCTCAGCCTCAAAGCCCCCTGTTCTTTA
 TCTGTGTGCTGGGGTTGTTGTAAGAGTTAAGTGCAATACACAGAGAGAGA
 GAGAGTACCTAACATGTATTATGTGCTCAGTCAATATGCATCATAGTACT
 CATTGTTACATATGTTCTTAAGTGCTTTATACGTTTTTTCCCTAAGTTGA
 CCATCTGTTTTTTGGCATTATGAAACATAATGATCCTAACAAATTAAAATT
 AAAAACATAAAGAATATTTGCCCAAAAAAATAAAGAACATGAATTCCTC
 AAGTAGCCAAAGGGGCCATAGACAGAAGTAAGCCCTTGGTGGGGCTTAGTT
 GAGAGAAGTCTCCAGAAGGTCTTTCGTGTGTTAAAGAAGAGGGTAACAGG
 GAGGAGGTGGGGAGAGATGTTAACTGAGTCTAAATGAGCACCTGGAAGAA
 GAGATGGGACAGGCCACTTCTGCCTGGACTCCCTGATTGTTAAGAAGAAT
 GAAAAAGAGCAGAAGTCTTCCCTGAGCCCAACTTCACTCCCTGACTTAAC
 CTAGTCTTTGCCCTTCCCTCTCACTCATGGCTACTTTCTGTGGTCACT
 TGTTGTAGAAATGGATGTGCAGCCACCTCATCTTTTTCTACCTCCTTCAC
 ATGTTTTAGATAATTTAATGTAGTAGAAGACGGTTACAGCAAAAAATTAC
 AAAAATCAAAATATCTCTGCTATCTACTGTTGCATTTCTAACCATCCCAA
 AACAGTAGCTGAAAACAGCACTCGTGGTCGAGCGCGGTGACTCATGCCTT
 TAATTCAGATACTCCGGAGGCTGAGGCAAGAGAATCACTTGAACCCGGA
 AGGTGGAGTTGTCAGTGACTCAAGATCATGCCACTGCACTCCAGCCTGGG
 TGACACAGTGAGACTCCGTCTCAAAAAAAAAAAAAAAAAAAGCACTCGTG
 TATTTTGTTCAGATCTGTGGTTTGGGCAGGGCAGGGCTCAATGAGGACA
 TCTCGTCTCCGTTCCCGCAGTGTGAGGAAGTGTAAGTGAAGTGGAGGGT
 CACACAGAAGATGGCTCCCTCAAGTGGCCAGCAAATTGGTGTCTACAATT
 GACAGGGAGCTGTTGACCAAGGGCCCCAATTCCTCTTCCTATGGCCCCCT
 CTCGGGCTGCATGGGCTTCTTTACAGAAATGGCAGCTGGATTCCAAGAGCA
 AGTATCAACAACCTACAGAAGAGTGGAGGAATATTGAAAGTTCACAGTCTC
 TTAAGACGTTGGCCAGAACTGGCAAAAGCTTCATTTCTGCCATGTTCT
 ATTGATCAGTCACAGAACCTGCACCAATTCAAGAGGAGAACATATAGAGG
 ACATCTCTCAATGGGATAAGTGTCAACAAATTTGCATCTATCACAATCTG
 TCTTTTGGGTACAACTATTTCTATTCTCTCATTATGCAAAATATACTCA
 CAACCTCCCAAGGGGTGCAAAAGCCTCATCCATTTATGGCAAATGTGGCC
 CTTTTAATTTATATAAAATAATTTGCGGGGGCTTCTTTATATTTTTAAC
 TCCCCCTGC

>Contig35

GTGCAGAGAAGTGATTTAAAGCCCTTCAGAAAGAATGCTTTATTCCCGTG
 GAATTTGGTAACCTTGCTTGGGTGTGGGGAGGTTTGTGAGCTTTCTCCACT
 CAAATTATCAGACCCTTTCATTTAGTGGTAGACCATTTCCCTCGTCCAG
 GCCAAGGGCACATAGTACAGAGAAATAGGGAGTTGTTACCCAGGGAGAGA
 ACTTGGCTCTAAACCTGTAATAGAAAGGTCAGTTCTGGTCTGGAGGGTCA
 ATTTTGATCTTTGGCTCAGATCCAGGAATTGGAACCAAGGCTTTTGAACA
 TTTTAATGCAGGGGATTAAAAAATGATACGAGTCATTACGAATATATT
 TGCTTAACATCTAAAGAGATCCCTCAAAACACTAGAAAAAATAAGAACAA
 AAATCTAATAAAACAAAAATTTGTTAAACACATTTACCAAATTTTTTTTTT
 TGGTAAAAATCAAATGTCATAAATAAAGCTAAAGTTCCTCTTGATGACT
 CGCTCCTCTGCCCTATTCCACTCCAAGTAACCACTATTATCAGTCTTGCC
 AATACCCCTCCAGACCTCTCTACCTCTATATACCATTAGAAGCACATGGT
 TTTGCATTGAGGATGTGCAGTGTGTTTGTGTTTACGTAATGTTATCACTCT
 GTTCTTGTTCATAATTTGCCCTTTTTCTCTCAATGATTGCTTGGCTATC
 TTTCTATTTCAGTAGCATCTCCTTTCTTTTAACTTACCATTGTTTATTT
 AACCTTGGCTCTATCAACAGATATGTAGGTTGTTTCTAGTTGATTTTATT
 AAGTATTTATAAACCAACGCATCAGTAGATGTCCATAAATTTCTTTACGGA
 AGATGGCAAGTAGTGGAATTGCTGAGCCAAAGAACATGTTTAAAAAACCC
 AAAAAAAGTAGACGCTACCAATTTTCTCTCCAAATGGCCATACCCACTT
 ACCCATACAGAGATGATTTGGAATCTGGCTTCTCACAAGGTGAGATGCC
 TTCACAGTTTCATTCTTCTGGCATGTCTCCCTTTTGTATCTGAGAGAG
 CTGGCAGAATTGTGCTCAATAATCAAGGATAGAGGGTCAAATGACAGCTC

FIG. 4 (13 of 61)

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AAGCTCACAGGCACCTCTGCTTTCTTCLAGACCACCTGCTTTCTGCLA
CCAGCTCTGTTCCATCTTATAGAATGGTTGCCACTTGGGTGTCTGCTCCG
ACAGCCATGTCTATCCTTTGCACTGCAGTTATGAAGCAGACAGAGCTAGGA
GAGGGGCTTTGCCAGCCTCTGCCCTAGCTTGGAGAACTTCAAAAAAGGAG
GGTATTGAAGTTGAACTCCCCCAAAAGGGGTGGTCCCCACACCTCAAAA
AGTGGTGCCTCCGAAAGAAATGTAAATTCGTGTGGGGGGGAAAAAGGT
TATTTAGAAATTGTTGGCTTGTCTGTGCCGAAAGTATGTGTGGTTACGGGG
AGTACGGAAATTTGAGGGGTGGGGGCGAGGCCGTGTGTCTTTAGCCCCG
GGGTTTTCCCGTCGCATGTTTAAGGGGGGGGAAGAGGGGGGATGTTTTCT
TTCCGCGAAGGTTTTTGAAGAACGGCGTGG

>Contig36

CCCCCACCAGCCACTACTCAACCGGCCGTTACGAAACAACCTCGCCACAT
CCACTAACCCGCTGGCTCACCACCCACCGCCCTCCCGATCCCCCAATCC
AAACTCAACCCCAACCAAGCGCCTCCCCCTCCCCACCCCTCCAGCT
CAGCCCCAACCTACCACCAACCCCGACTCGCCACCGAAAAACCAACAGCA
AACCCAAATGCCACAAAACCAAGTGTCCAAACCTCCTTCCCATCAGTTT
GGTGGGCCCATCACCCTTCCCCCTGGCCCAGGCTCTCCTTTTGTGCGCTT
GGAGCAGCAGACTGATCTCCAGCCTTCACTCACTTCATGTGGTAATCTG
TTGTGTTTCACTGTGCAATCTTCTGCATCCCCCTCACTACTCTGCTGA
AAACACTCTAGTGGTCTCTATTGCTCATTATGAAAGTCTAGATATTAA
ACGTAGAAGGCCCAGCACAATTTGCCCTATGCCACCTACCTCTCTAATC
TTTTCTCCTTACTCTGACAGACTCTCCGTCTGTCTATTATGTATTCTTTT
ATTGCTCTCTTCTACTTTTAGTATGAACTGGATTATGGATTTTTAAAC
ATTGCTTTCAAGTATGGAATAAAGAATTTTATTATTATTATTATTATT
ATTTGAGACTGGGTCTCACTCTGTTGCCAGGCCAGAATGCAATGGTGCA
GTCATATCTCACTGTAACCTCGAATTCCTAGGCTCAAGCCATCCTCCTGC
CTCAGCCTCCTAAGTAGCTATGACTACGGGTGTGCATCACCACATCTGGC
TAATGGAATAAAATATTACAATGCCTAATCTTAATTTTCAAAATTTTAA
TTACATTGTACCTAATGCCCATGCATTACTTTTTTTCAGTGGGTCAATAG
CCCTCACTTTGGCAAAGGTCCCAGGCCCAAGGTAAGGCCCTTACTTTTTCC
AAACTCATCTTTGAAAGACATAAGTGCCTGTAAGTTGTACCACATTAGG
TTCTAGGAATTTTTCATCAAAGACTTTATCAGACTATTTTCTCTAAGTT
GAGAAAGAGCTGGGGGCAGAATATGGCACTGAATGACTGAAGAGAAGGCA
CTGAAATCAGGCCAGAGGTTGCTGGAAAGAGCAATGAGGAACACCAGCAG
CAATGAGGAGCCGGTGATGATTTTGGCTTACAGGGAGGTGTGTACCACA
CCGATTTTATCTCTACGTGGATGAACCACAGCTGTGCGCTCCCTTGTCTC
CAGGACATCACACTCTCCACATTCCCTCCCATCTTCCGGCTTCTGCTTCC
CGGGGCCCTCATCTGCCCATCCTGGGTGAACACTGGTGGTCAACTGCT
GGGCGTACCTTCCCGCTCTGCACACCCTCCCTGGCCACCCCAACCACTCT
CACGGCTCGCACTGCAGAGGAGCCGCATCTCTAGCTCCAGCCCATCTGCC
TCTTCTGAGCTCTAACTTCACTGATAGGCGACTCCTGCCGGTGTGCTCAC
AGGCCCATCATACTTCAAAGCATTTTCCCCCTCAGAACACCATGTCTGGC
TGCTCCCTCAGAAGATACATCTCTCAAGCACATCCCCGCGCTCTCACC
TGGATGACTGCATTACCTTCTCCACATTGCCCCCTCCTTGGATGTA
TATAGATTGTTTTAAATACAAATCTGATGTGCTTGTCTCCTGCTTGAA
ACACCTCAAACTGCCTTCAAGATAAACCACTGCCCTTGACATGTTTACA
GGTTGCCCATGGCCTGGCCCTGCCATCTCTTCAAGCTCATCTCATGCCC
CTTGCCCCCTCGCTCTCTGGGCTTCTGCTCCCTAGCCCTCCTTAGGTTT
TCTAACACACCATAGTCTTCTAGTGTGGGGCTCTGCAAGTGTCTGTTT
CCATTGCTTGAGACATGAATCCCTCTCCCTATCTCTACCTGCACCTTCAT
CTGATTAATCCCTACCTTCTCTACTCATGATGTTGCTTTCTCAGGGACTC
TCTCTGACTTTTAACTAATCAGGGTCTCCCCAGTATATATCTTCATAG
CACTCTGTATTACTCTTTTCTTAATGACCACCTGCTGTAGACAGAATGTT
TGTCTTCTCCAAATCATATGTAAACCTTCCACCAGAGCGATGATTAG
AGAAGCCTCCC

>Contig37

GACTGACATTCAGAAGATATTAATAAGAGCACTAATGATGGGGATTGCAA
CCATGTCTTTACTGACTTCCAGAAGCTTCTTACAGTAAACATGAAATCAC
ATAATTTCTTCACTTTCTACTGTTTCTTGTCTGGGCTCTGTCTCTGCT
TACTGTCTAATATCTTGGCCCCCTTAAAGTTGCTAATCTTCAAACCTCA

FIG. 4 (14 of 61)

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TTCTGTGACTGGGCCGCTGGTCCTTG...ATGGGCCTTGAAGATAC10A
 CTGTACACTTATCTGGAGCATCCAGTGCCTACCACCTGACCCAGATTCCCT
 CATTGCGCTCCTCCCTCCTCCACCTAATGGGATTGCTCATACCCGTGTG
 GGACCCCTCCCATTTTCCCCAACTGAATACTTATCAAGACAACGCATTGC
 CATACTCCCTCGTACCCTGCTCTGGGCATCAGACTGAATGTTTGTTCCTA
 TTGAGGATCTGCAGCTGCATCAGTTTCCCCAGCACCGTCCAACCCCTTGA
 GCATGGCTAGTCCTAAAGCAGAGAATTAGCCTTTCTATCCCTGCTGCTAT
 ACATGCTGGGACAAATAATAAGAAATGACAGCATTTTATGATAATGCAGG
 CTGCAGGAGGCAGGAGGCAGGAATCAAATTCGTGCTTATCAAATAGTGCT
 CCAATTCCTTGAAATATTGGACTATAGAATATGTATGGATCTATGCTCAG
 GTGGGTTCCTATTACTCACTCCACTGAGGCCAGGTTGTGGGATTAGCTG
 TCCAAGAGGGAGTTTCAGTCTCACAGCATAGGGTCATTCTGAGAATTACT
 GGCCCACTTGTGTGGAGACCTCCAGAGAACAGAATCTGGGTTGGTGCC
 ATGTACTTCCAGGAGGAGAGAAGTGGCAGGATGCCAGCCCCACAATCAG
 AGGGGAAGGGGCAGAGCCACATGTATGAAGATCCTCTCCCCAGTACGTGC
 CAATCACAGGGCTTCCTAGCTTTTGGGCCAAGGAAACAATGTGGGAAGCA
 AAAAAAGCAATTTTCTCCTCCCTTTGCATGAAGACTGAGCAGTTTACC
 AGATTCCTCAGGGAACACCCCTTCCACTCTGGGTTGAATGTGAGTGAGAGA
 CATTGAGCTGGAACACTAGAAAAACTATTTCTGAGCCACTCACCTTTAG
 CCTAGAAAAGTGTGGATTTGTCTTCTATCTTTGCCACAGTAGAGACTGC
 TGATAGCATCAGAACTTGGGCTCTGGAATTAGACAGATATGGGTACAAAT
 CTGAGCTCTCTCACTTATTAGTGTGGGATGTAGAGCAACTTTTAAATCC
 TTCCAAACCTCAGACTTCTCATGCATGATGTGAGGATTGTAATAGGGCCC
 ACCTAATAGGGGTTTTTGAGAATTAAAAAGTTATTCAATGAACAGCATT
 TAGCAAGATGCCTGACCATTGAGAAAATAACAAATTGTTTATTATTATTG
 TTATTATTAAAACTCTTTCTGCACCTTCTGACTGGGGGCATCGTATCAT
 CAGAAATACTTAGGATGGGATGGATTCTGCATGGGCTGAGTCAAGGGTG
 CAATAATGGAGGAGTGAAGAAGGAAGAAATGGAGGCAGAAATCCCCAGGA
 GCCCAGCATGGTACAGGCTGAGCTAGTGTGCAGAGCCTCCTTGGAACA
 GCCACAGAGCTTGCATCTGGCCCTGGAGGAACCTCTTCTAGCTGGCAGGA
 CCAGCCACAACAGTGGCCAGGGGATTTCCAGGGCGTGGGCTCCTCAGGA
 GTTCATTGTGACCAAGCCTGCCTGGAGAGGGTTATAACAGGGATCCTTC
 CCTACTGGCAGGTGATTTTACCCCTCGGTGAGAAGCTCAGGCATTGTTTG
 ATGGAAGGTGGAAGGCCCTGTGCTGGGCCAGTGACTATCAGGGATGGGCG
 GGTGGCTGGAATAAGCAATAAGACAATATGATAACACAGTTAACCACC
 AACTATGTGAAGCTACAATATGGGTATCTGTAATAGACAATTCCAATGT
 AGAGAATAATTTAAGGTGTCAATCTCCCCGCCAATGCCATAAGCACACG
 GCCTCTGCCTGGGTTTCTCACTGTGGAATGTCTCCTGGTCTCCTCATGC
 CCAGAGAGTGGGAAGTACTCCTACTTTAACACCGGCTTTCTGTCAATTC
 CNTGCAGCCCTCCTCAGCCCCCTCTGCACAGGGAGGTTTCTCCTGCTG
 CTGCAGTGCTTTGACTTTGTTAGTGGTACCTGCACACAGGTATTGGTGTG
 CTTGTCTCACCACCCTACATCACTGTAAGCTCCCCAGGAGCAGGCTTCCT
 GTTTGACTCACCTGTGATCCTCCACCTCCCACCTGTAGTGCCTCAAGCA
 TTCTGTAGAGCACATGGACGCC

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GACTAATAAGTACTTCATTATTTGGGTATTTTCCAAGAACAACATATTGT
 AGGAAACCATTTCTTCTAAAAAAAAGTGTCTTTTAAAAAGGTGAATA
 ATTTTGTCTAATTCAAAGTTTATTGAAAAGTTATGTATAAAACAAGGTA
 AAAGGAACAAGGAAATAAGGGAAATGTAAAGAAAATTATAGAAAATAAAGT
 GGTATTTTTTGGTAAGAAAGCTTAAAGAGAAATAATTTTAGGTAAGAAAG
 AATCTTACCTAAAAATTTGTGCTAGAATAAAGTGACTGGCTAAGAAAGGG
 ATGTTCAAAGCTATTTATGACAAACCCACAGCCAATATCATACTGAATGG
 GCAAAAGCTGGAACATTCCCTTTGAGAACTGGCACAAGACAAGGATGTC
 CTCTCTCACCCTCCTATTCAACATAGTATCGGAAGTTCTGGCCAGGGCA
 ATCAAGCAAGAGAAAGAAATAAAGGGTATTCAAATAGGAAGAGAGGAAGT
 CAAATTTTCTCCGTTTTCAGATGCATGATTGCATATTTAGAAAACCCCAT
 CATTTCAGCCCCAAAACCTCCTTAAGCTGATAAGCAACTTCAGCAAAGTCT
 CAGGATACAAAATCAATGTGCAAAAATCACAGGCATTCTATACCCAAT
 AATAGACTAACAGAGAGCCAAATCATGAGTGAAGTCCCATTCACAATTGC
 TACAAAGAGAATAAAATACCTGGGAATACAACCTTACAATGGACATGAAAG

FIG. 4 (15 of 61)

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ACCTTTTCAGGGTGAA...GCAAACCAC...CTCAAGGAAATAAGAGAG...A
ACAAACAAATGGAAAAACATTCCATGCTTATGGATAGGAAGAATCAATAT
CGTGAAATGGCCATACTGCCCAAGTAATTTATAGATTCAATGCTATCCC
CATCAAGCTACCATTGACTTTCTTACAGAATTAGAAAAAACTAATAGCC
AAGACAATCCTAAGCAAAAAGAACAAAGCTGGAGGCATTGTGCTACCTGA
CTTCAAACCTATACTACAAGGCTGCAGTAACCAAAACAGCATGGTACTGGT
ACCAAAACAGATATATAGACCAAAAGAACAGAACAGAGGCCTCAGATATA
ACACCACACATCTACAACCATCTGATCTTTGACAAACCTAACAAAAATAA
GCAATGGGGAAAATAATTCCCTATTTAATAAATGATGTTGGGAAAACCTGG
TTAGCCATATGCTGAAAACCTGAACTGGACCCCTTCCTTACAACCTTATAC
AAAAATCAACTCAAGATGGATTAAAGATTTAAACATGGCTGGGCATGGTG
GCTCACGCCTGTAATCCCAGCACTTTGGGAGGCCGAGATGGGTGGATCAT
GAGGTGAGGAGATGGAGACCATCCTGACTAACACAGTGAAACCCCTGTCTC
TACTAAAAAATACAAAAAATTAGCTGGGCATGGTGGTGGGCGCCTGTAAT
CCCAGCTACTTGGGAAGCTAAGGCAGGAGAATGGTGTGAACCCAGGAAGT
GGAGGTTGAGTGAGGCAAGATCAGGCCACTGCACTCTAGCCTGGGCAAC
AGAGTGAGACTCCATCTCAATAAATAAATAAATATGGAACCTCTCCAACA
CAATAATAAGACAAACCCCCAAATGTTTTAAATGGGCAAAAATATTTGAA
CAGACACTTCACAAAAGAGGATATGTAAATGGTCAAAAAGCACATGAAAA
GATGTTCAACACCATTGGTCATCAGGGCAAGAAAACCTAGAACCAATG
AGATGCCTCTGTACACCCTTAAATGTCCAAATTAAAGAAAAACAAGTTT
GGCAAGTTGTGGAGCAACTGAAATGCTCGTGATTGCTGGTAGAAAAAC
AAAAATGGCATAACCATCGCAGATAATTTGTTGTGAGTTTCTTACAAAGT
AAACATATACTTTATGATATGACAGTTCATTCCAAGAGAAATGAAAACA
TAAGTCCACACAAAGACTTGTACCTGGGTGTTTATGGTAGCTCTATTCAT
AATTGCCAAATCTGGAAACAAATCAAATGTCCATCAGCAATGGAATGGA
TATACAAATTGTGGTACACATGTACAATAGAAAACCTACTCTGCAATGGAG
AGAAATTAACCATTGACAAACACAAAAACATGGACAAACCTCAAAAACAT
TATGCTGAGCAAAAGAAGCCAGACACAAAAGACTGCTCAGCGCATGATT
CATTATATGAAATCACAGAAAGGTCAGTTGAAGGTGCAGAGACAAAAA
GTAGATCTGCAGTTGCCTGGGGATGGGGTGGGAGGTTGACTGCTCTGACG
CGTAAGGAAATTTGGGGTAGGTGGGGGATGGTGGGAATATTTTTTGAAT
TGAATTTGGGTAATAGTTTTAATAGGTAAAATATTGGACCCACAGTATT
GAGATAGGTTTTAGTCAATTTAGACAGTTTATTTTGCCAAGGTTAAGGAT
GCATCCGTGACCCAGCCTCAGGAGGTCCTGACAACCTGTGCTGAAGGCAG
TCAACATACAGCTTGCTTTTATTCATCTTAGGGAGACATAATACATCAAT
CAATGCATGTAAGGTTTACATTGGTTCAATCTGGAAGGTGAGGGAACTT
AAGTCTGGGTTATGATAAGCGGTTGTGGAGACCAAGGTTTTATCTTGTAG
ATGAAGCCTCCGGGTAGCAAGCTTCAGAGGGAATAGATTGTCAAAGTTTC
CTATCAGACATAAGGTCTGTGTTGATGTTAATGCTGGTCAGCTTTTCTG
AATTCCAAAAGGGAGAAAGGTTACTGGGGCATGTCCAACCTTCCCTTCC
ATCATGACCTGAACTAGTTTTTTTTCAGGTTAACTTTGGAATGCTCTTGGCC
AAGAAGAGGGGTCCATTGAGATGGTTGGGGGGCTTAGAATTTTATTTT
GGTTTACAGTGAAGACTTTTCAAGCTAGACACTTAAATGAGTATGTTGCA
AAATGGCAATTTCTTAGCACGGC

>Contig39

GACGTCCTAAAGAAATGCTAAGGTAACCTCAATTAACCTATGCTAGAAAAGA
GAGTTAAGTATTTAGGAGGATTTAATATGGTGTAAAGTTGTGAAAATCA
AAATGGAGACACTAATGTTAAGAAAACCTGATAAATGGAGCCAGGGAAG
GCCATGAAGAAAGAGTTCTCACTTGTATCCCTGATCATGAAAAAGACT
CTGCAAAAAACAAAACCTTGACAAAGGCCATTGCAACCTTACACAAAAA
ATACTACTTTAAAGGACATGTGCCAGCAACTGCCTGTCCAACCTCAGA
CTGGCAATATCTTTGTTATTGATCTTAGTAGCCAGCATAACTATTTCAA
AACAGTGATGTAATGCTCATTTTTTTTCTTTTGAAAACCTTTGTCTTCT
GTAAAAACCTTTGTCTTCTTACTTACCCTGAATATGCACAGAGTTTACT
ATGGAGTGCAATTCCTGTTGCAATGCTCTATTCCCAACAAACATCATT
TTCTTTTAGAGAGCCTCTCTCTGTTTGTGATTTAGGTTGGTGATGTAAAG
CAATGGCATAACTGAACACTGATTCAAAGAAAAGTGGCTTTTCTCTTTGT
TGATTAAAAAGAGGCCTTATAAATAGGATAGTAAGATTTGTAAGTTGAA

FIG. 4 (16 of 61)

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CTTAAAGCATGAAGAAAATTTAGGGGCCAGGCAGGGTGGCTCACACCTGT
AATCCCAGCACTTTGGGAGGCCAAGACAGGAGGATTGCTTGAGCCAGGA
GTTCAAGACCAGTCTGGTCAACACAGACCTCATCTTTACTAAAAATAAAA
AAATTAGGCCAGGTGCAGTGGCTCATGCCTGTAATCCCAGCACTTTGGGA
GGCCAAGGCGGGAGGATCACTTGAGGTGAGGAGTTCGTGACCAGCCTGGT
CAACACGATGAAACCCCATCTCTACTAAAAATACAAAAAAATTAGCTGGG
TGTGGTGGCGGGCACCTGCAATCCCAGCTACTCGGGAGGCTTCAGGCAGG
GGAATCACTTGAACCTGGGAGGCGGACATTGCAGTGAGCTGAGATAGTCC
CACTGCACTCCAGCCTGGGCGACTCAGCAAGACTCTGCCTCAAAAAAAA
AAAAAAATTAGTCAGGTGTGGTAGCACACAGCTGTGGTCCCAGCTACTC
GGGAGGCTGAGGTGGGAGGATCATCTGAGCCAGGAGGTCAAGGCTGCGG
TAAGAGCTGAGATTGTACTACTGCATTCCAGCAGGGGCTACAAAGTGAGA
CCCTGTCTCAAAAAAGAAAAAGAAAAAGAAATTATGTTTTTAAATTTA
TAATTATAATAAATTTAATTACATAAATTTAAGCTCAAGTAATTGTAAAT
ATTCTTTCTGTGCACATAAGTTATTCTTGATTGACCCACAGGAGCTGG
CCATTCTTCAAGTCAGAAGGCCTGAGAGAGGAGCTGCCAGSTGGTCTTC
ATGGGGCTGTGCGGCCAGTCATCCCCACAGGTTGACAATCCTTGTTGAC
TTCATCCTCGTTGGATCCTCTGTATCCCTGACGATGAGCAACTGTGAGGC
CCGTTTCAGCACTGAGTTCCAGTCAGGAAAAATCCACCCACCCACCACA
CGCTCACACTTACACACACATTACACATGCACACACGTTCTGGCTCCGA
AAAAGAAAAAAGCAATTTAAATAATTCTGATCCTTTGCTTATTT
CCACAAACTCCATGAAAATTGTACATTGTCCAAGCAACATTTCTTAATAT
TCTCTTTTCTCTCATATCCATTTTCTTACTGCTGTCTCCACCTTTCTC
TTCCAAACTCCCTGTTAAATCCCTGCCCGAGCAACTTTTATTCAATTT
TGTGGAATGGAGGCTGCTCTGATTTAAATTAAAAAATAAATAAATCCC
TACTCCATGTCCCAGATCCCTAGTTGTTTTTGTGTTTTTGTGTTTTCTGAG
ACAGGGTCTTGTTCTTCCATGCTGGAGTGCAGTGGCATGATCATGGCTC
ACTGCAGCCTCAACCTCCTGGGCTCAAGTAAATCTCTTGCGTCAGCCCTC
CCCAGTAGCTGGGAGTTCAGGTATGTGCTACCATGCCTAGCTAATTTTTT
TCTTTTATTTTGTAGAGACACGGTCTTGCCAGGTTGCCAGGCTGGTATA
GAACCCCTGGGCTTAAGTGATCCTCCTGCTCGGCTTCCAAAGTGCTGG
GATTACAAGTGTGAGGCACTGCACCCAGGCTGGATCCCTGCATTTTACA
GATTTAGCATCACAAAGTCTAAACAATTAGACTGACTAAGGCAGAACTG
CCCTTATGACAGCAGACATAAGAAGGAAAAGGCCAAAACACTGTGTTAAA
AATTATCCAAATGTGAGGAAAAGGCCAAAGAGAGTAGGTGTGCCTTTTGTAG
TGTCTAAGCTGCCTGCCAAGGGGCATCTGATGCTCTCAGGCAGGAGTCC
ACAAATTTTTTTTGTAAAAGATCAGATAGTAAATCTTTTCAGCGTGAAG
AGCATGAGTCTCTGTCAAAATACTCAACCACCATTAACATGAAAGC
AGCCAACAGACAACATGACAAATGAGTGTGGCTGTGTTCCAGTAAATC
TTGATTACAAAAACAGGCAAGAGGCCAGAGCTGACCCATGGGCCATAGTT
TGCTGACCCCTTCTGTAAAGGAAAGTATTTTTGTTTGACTTGCTGTTTAC
CATTGATTGAACACAAGGCTCTGTAGAGTTACTTGTTAACTGCAGAAGA
TTGATGAGTGGCAAGTAATTTTATTACCCAGAAATATANNATTATTCTGT
TCAGTAGATAAGATAAACCCACTGTTATATTACTGTCTTGTTTAGAATGT
GACTTTGATTCATTTTTTCACAAATTCATATTATTGCCCTAATTTGTATA
TAAGTATGCTTCTTTTAAAAATATATATTTTTTAATAAATTTGAGACAGG
GTCTCACTAGGTTGCCAGCCTTTTGCTATAATGAGAGCATAAAGTGAAT
TTCACACTTTAGCCTAGTGCATAGATGGGATTACAGGCACAAACCACTGC
ATGCAGCTAACTTTGCTTCTCATTCCAGCACGTTCTATTCCNNNGNTTTT
CATATACGCGTCTCTTAATGC

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CGCATTAGCCCAAGTTTTCTTCAGTGTTAAGGTTTTTGTACTCTGTGC
CCAAATGTCTTCCAAAAAGGTTAAGTTTTTTACCTTCTGCCAACATT
ATATGAAAGTGTCCACTTTTGTAGACTTTTACCAATGCTGACTACTTTT
GTTTCAAAAAAGCTCTCAGTAATTTTCTATTAATTACTTTTACCCTTTT
TATTGAGGGTGTTCACCTTTTTATTGTTAGCATATTCTCTGCGGCTCCA
TTGGACGCTTGGCAGCTTTTTGGTAGTAGGTGCCTTTAGAAAAGTCCTT
CTCGTCTGGCCCTTTCTGAGCAAATCTAGTGAACAGAATTGGCTCCATGC
TCAGCATTGCTTAATACGGTTGATCCAGGGCCTAGGACTCATTCTTCAT
TACCATCCACTTGCATTGTCTTAAAGCAAGGCTCTATTAATTTAATTTGG

FIG. 4 (17 of 61)

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CATTTCCGTGTCAGCTCTTAGTTCATTAAACAAAGGCTTTAGAAAAC
TCCAGTAGATGCCTATGTTGCTTCCTTTTAAAAAATTTTGGAGCTGTTT
CCCTAGCCTAACCTTTTCTTCAGGGCAGGAGTTAAGTCCCTTCTACTGCA
TTCCCTGTGAAGATGGTGATTCAAGAGGCAGGGCACCTGTTGCTTTGTGAA
ACAGTCCACTCTGCAGCTGGGCAGCTCTGTTACTAGAATGTTCTCCCTTC
TGGGGAGCCAAATATTTTGATGTCTCTGTGAATCTCATCTGCTTATCCCA
TCGTGTTTATGTCTTGAAGATGCACAGGTCTGACACCACGAGGTAGCCCT
TAGAAAATTTGATGGCATTCTGATGTGTCCCAACTCTTCTCCAACCACT
CCTCCCAGAGCTTGTTTCTTAAGCCCCCTTGTGGAGCTGATTGCTTTCCTC
AAGGCAGCTCAGTTTTTCCCAGTTTGCTCCTGGTGGTCTGAAATATGAT
TGACTCCTGAATACTCCAGGTGTGAAGGAGAGTGGGGTGGCCTTTCTAC
TTGTCTATGGCCTGGGTTTTAAGTTGCTGTCCAGTGGAGCAGAGGTGACTT
TCCCAGTGAACATACATTTTTTCCCCTCTAAATCCTTAGCAATTTGTCTC
CAGAGGCAAGACCTGGCCAAACCATTTGTGTTGAGGATTGAATCAAGAAT
GATTGAGGAGATGACAGTAGTCCCCCTCATCTGAGGAGGGCGTGTCCA
AGCCCCCTCAGTGAATGCCTGAAACTGTGGATAGTACCCAACCTTATATGT
CTATGATTTTCTATAAATTAATACATGCCTGTGACAATGTTTAAATTTAT
AAATTAGGCAAAGAGGCCAGGCGCAGTGGCTCAAGCCTGTAATCCCAGCA
CTTTAGGAGGCTGAGGCCTCACCTGAGGTGAGGAGTTCGAGACCAGCCTG
ACCAACATGGAGAAACCCCGCCTCTACTAAAAATACAAAATTAGCTGGGC
ATGGTGGCAGGCGCCTGTAATCCCAGCTACTCGGGAGGCTGAGGCAGGAG
AATCACTTGAACCCGGGAGGCGGGATTGCGGTGAGCTGAGATCGTCTCA
TTGCACTACAGCCTGGGCAACAAGAGTGAAACTCCGACTCAAAAAA
AAAAAATTAGGCAAAGAAAGAAATTAACAACAATAAGTAATGAAATAGA
ACAATTTCTAACAATATACTATAATAAAAGTTGTATGAATGTGGTCTCTTT
CTCAAAATTACCTTTTTTTTTTGGAGACAGGCTCTCACTTTATTGCCCAGG
CTGGAGTGCAGTGGCAGCATCACAGCTTACTGCTGCCTCGACCTCCTGGG
ACCAAGTGATCCTCCACTTTAGCCTCCTGAGTAGCTGGGACCACAGGCA
TGCACCACTGTATCTGGATAAATTTGTTTATTTTTTGCAGAGAGAGG
AGGTCTCACTATGTTTCCCAGGCTGGTTTTGAATGCCTGGGCCAAGGGA
TCCTCCTGCCTTGGCCTCCCAAAGTATTGGGATTACAAGCGTGAGCCACC
ATGCCTGCCCCAAATATCTTATTGTTCTATACCCACTCTTCTTCTGT
GATGATGTGAGGTGATCCATTGCCTCCTTGATGAGATGAAGTGAGGTGAC
TGATGTGGGCATAGTGATGCAGTGTTTAGGCTGATATTGGCCTGATGATA
TGTCAGAAGGAGGGTCATCTGCTTCGGTGATCCTGGATCATAGAGTCATG
ATGATGTCAATGGTTGGATGTGAGGAGCAGACGATGTCAATGACTAACGA
TAAGCTGGACAGGTGGGATGGTGGCACAAGATTTTATCACGCTACTCAGA
ATGGAGCACAATTTAAACTTCTGAATTGTTTATTTTTGGAATTTTTCAT
TAATATTTTGGATTGCAGTTGACTGTGGGTAACTGAAACTGTGGAATGT
GAGACTGTGGAAGAGTGAGGGAGTACTGTATTATGGAACGTGAACCTCTAT
TCGGTAGGGGAACAGAAATTCACATTTGTGGGGCCAGGTCTCTGCATCTG
TAGGGATCCAATTGTTTCATTTCTCGTTGTAGCAAAACTTGGCTTTGGA
ATCAGACAGATTGATGTTTGCTATCATTCTAAATGGGTGCAGCTACACTT
TCCTCAAGAGGTAGTTCTGAAAATTTAACAATAATGTGAATTTCTGGTAA
AAAAAATAACCTCAAAATATTAGTTTTCTTTCTTTGTGTCTGATGT
ACTCCATCAAATACTGGGAAATATGTGTCTCTCATAGAAATGTCATGGAT
CTTTGTAATCTGATTATCCACAAACCTTGGGGATTAGCTGTTTCAATGT
TCCTATTTTACAGATAAGAAAATGGAGCCTGTGGTAAGTTAAGTGAGTTA
CTCATGGCTACTTAACTAATATTTTACTAGGTGATAGGCCAGAGCTAGAG
CCCAGGTACCTTCTTATCAATGCTCTGCCTTGTCTGTGCTTCTGT
CTGTCTGTATGTGTATGTGCTGTGACAGTAAGGCATAGTTTAAACCCAG
TAGAACTACCGGTTTGTAAATGAATTCACCTGTAAATGACTGACCATTCA
AGGAACAAGTGTTTTTCTATGCTTGACACCTGTTTTGGATGCCAAAAG
GATACAAATGTAACCTCAGACACTCTGGGCCTCATTTTGCACCTATTAGC
ATGTCCAAAATTAAGAAAGACTGACCACACCAAATATTGGTGAGGATGTGG
AAGAACGGGAACCTTTCATACACTGCTGGTGGGGATGTAAATGGTACAAT
CCCTTTGGGTAAACAGTTTGACAGTTTCTTAAAAAGTTAGACATATATATT
TACCATATGACTCAGCCCTTCCACTTCTAGGTCTTTACCCAAGAGAAATG
AAATGCTGTGCTTTTACAAATGTCTATACAGGAATGTACATAGCAACCTT
ATTTGTCAATGCAAAAAACAGAGACAATTCAACGTTGTCAAGAGTGAATG

FIG. 4 (18 of 61)

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GATGAGCAAGCTGTGGT CTCTATGCA...GGTATCCTACTCAGCCAG
AAAGATATGGCTAAT
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GACAACAATGTCATGCATAAGATGACGATGGCCTGGGTGATTGATGCAAA
CAAGGATAAAGAAAAATAATCAATTTTGTCCCATTTTCAAAGACAGATAG
CAGCAGCAAGAGTGTAAAGTCTGAGGAAAGTCATATTCCTTCCTCCTACAA
CATAGCACACACACTTACAAAAACAATACACAGACTCCTGGCCAATGGAC
TTCAAAACTGAGGAGGATCATTAAATTTAAATGTTCCACCGCTGCATGAAA
TCTCCCTGGGTCTGCCCCTCCCCTCCCCACCCTCCTCCACTTGGGCCGGG
GCACAGCAGTGATTCTCTCACCTCTCAGAGTGAGCCAGTGTTGGCTGCAT
TGAAGGCTCCAGATATGCAAAACAGGGCAGATATTCCTGGACCAGGGTGCA
CAGAGTGAGGCTCCAACGCACCCTATTAAGTGCATGAAGGATGAATGAGC
CTCTGGTATGGGCTGGGACAGAAAAAGGATTCAAGGGGCCCAAAAGGGT
TTGGGTGGAACCTACCAGGAGCGGCAGTACAGACTCCTTGGGAAGGTGGC
CATGATTTAGCCACATTACCAATAGGATAATCTGGAGAATTTCTAGCT
TGAGTTTCTGGGAGAAAGCAGATTTCTGGATTATCTGGTGACAGGTAACA
GGGCCGAGTTCATCCACAGCCACCTGCAGTGTTAGCACCTTAAGCTGAGT
TCCTTGCCACAGGATGCTGTACGCCCAGTCAGTGAGACGGTTCTTGG
CTGAAGGACTGAAAAGCTTGGGTAAGTGAAGTTCACCTAAGCCTCTATCTC
TTGCTCCCGTAAGTCAAGGCTCATTGTGGCTCCTTGACGGCTTGACTTCA
GGGTTAACAGAGAAAATGAAGGTACAAGTGCCTTGTGAAGTCTGAAACTC
CAAACAGTCATTCTCAAAGTGCCGTCCACAGTCTAGCACATCAGCATC
ACTGGAAGCTTGTGTTGAAATGTAAATTATCAGGTCCTCCAGAGCTATGTA
TGAATTAGAAACTCTGGGAATGGGGCCCTGCAATCTATTTCAACAGGTCC
TCCAGGTGATTCTGATGCAAGTTAAAGCCTGAGAACTCTGTCTATACA
AATGGATGTCAACTCAAGCTGCTCTTCAGAATCACCTATAGCACTTGTTT
ACCCGAATCCCTGAGAATGGAGCTTCAGGACTGCTATTTCTCAAAGTTTG
CCTGGTGATCCTGAGATGGGGTTTGGGGGACAGAGATCCAAGGTGCTACC
AGGTGTGAGGAATTGTTAGAAGGCAAACCTGGCTGTCTAGGGTGCTT
AAAGGGTACAGATCCTAGGATTCTGCCTCTTACAGCTGAATCAGACTTTC
CTAGAATGGGATTGCTGTCCAATGGCATGCCTCCTGGGTGACTCTGATGT
ATAGCCTGGGCTGGGAACCAAGAGGATTATCTTCCATTGACCAAGCTG
ACAAACTCGCTTAAGGCTCTGAGTTTCACACTTGATTTTCTAGCCCCTGT
CCTTCCATGGATCACCTGCCCCCTTCCCTCCTAATCAGGAGCACAGTCAG
TGGATGCACCTAATGTGGCCTCTCCTTGGCTGCAGGGAACAGGTGGAAATG
TGGCCATAGGTGTGAGGGCTGCCTGCCATGTATTAATAGCTACAGATT
GAAAGATCCAAGGACAAGAGACTAGAAAAAAATTTAAAAACAGCCAAGCAT
TGGCCCAGTAATGGCATTTCAGAAATCCACCAAAATATTAAGATGCTTTT
TGAAAAATATCCAGAGCACTCATGTAAAAGTGCTTAATTATTAATAAAG
CTGACATGTGTTGGGTACTTCTGTGGGTCTGGCACTAGGCTAATTATGT
TTTTAGGAGTTGACTCAAATGCTCCCTGTCTATAATTATGTGAAAAAATAT
AATTATTAGCTCCATGGTACAAATTAAGGAGAGGTTACATAAATAAAAAG
GAATGATACTCAAATTAGTAACCAAGAGCCCATGCTCTTAAACACTATGCT
ATTATTTGTGACTCTTACATAGGTGGCAAAAGTCAAAGGCTAGATTGAC
TTCTGTCCACTTCCAGCCAAGATGAAGTACAAGATTCAGATACACCCTTC
CGCATTAAACAACCTTAGGAATCAGACAAAATATACAAAGCATTGTTTGT
ACACATTGGATAACAGACAGCACTAGATAGTCGTGTCTGAGAAAAGCGGT
GAAATGAGCTGAGTCTTAGAATTGCCCCAGTTTACTAAGGGGCATAGTAA
GGGCATAGCTGCAGCACAAAGAAGCAGAACCCACAGAGACTGGCGTTCA
CCTGAGTTGAGAAAACCAAGTTGAAAATTTAGGAACACTAACACAGATAT
GTAGGCAAGAGTATCAGAGAGGAGACAGTTGTAGGGAAAAAGAGAGCTTT
ACAGAGAGACAGCGAGAGCTCCAGAGACCCGAGAAGATTGCCCTGACGT
CACTAGCTGAGTACCGATCAGTGCATACATGTAAGGATATTACTCAATAT
GTGGAAAAGAACAGAAGGAATGATGTCCAAAGCTCACCAGAGCAGGAA
TCATTTATGTTTCCACCAGCCAGAGTGGAACAACCTTGTAACGCATATGG
AGTACTCAAACGAATATTTCTCAATAATAAGTTCAAATTAAGTGAAGT
AAAGCCTGCCCCGCTTTGTCTGGACATGCCTAACAAAGCTTTGAGGGAAGC
CTCAAAAGAATGAAACCGTGTCCAAGTAATTTAAGTGTGTCCAGAAAAA
AATTCAAGAACATTTAAATAAATATTAATAATATGATCAAAACCCAGCAAGG
TTAAATTCAAATGTCTGGCATCCATTAAAAAATTACCAGCCTTGAAAAAT

FIG. 4 (19 of 61)

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TGGCGGGAAAAATATTA: .ATAATGAA .GAAAAAGCAATCAACAGAN
AGGCCTAGAAAGTATACATATGATAAAATTAGCAGACATTAAATGGTTAT
GATTAATTTATTTTATATGTTAAAGAAGGTAGAGAAGAGCATAAGCACAT
TAAAGAGAGACAGGAAAGTCCAGTACTCACACAGGGCCAGGAGCAGTTT
TCACCAGTCAGGTGGGAAAACCTTCATATTTTCATGGAGCATTGGTAGAGTA
CACAGTGTCTTGCCTTAGTAGAGGGATAAATGCTGTTCTGTTCCCGCCTA
ACCCATCTTGAAAGAAAATCTGAAAGGATCAAACCTGTATTCAAGTAACCT
AATCACATCCAGCACACAGCTCGACTAGTTATAAAAAACAAAAATATTA
ATATCTAGAAACACAAAAATAATATCTAGCACCCAACAAGGTAAAATTCA
CAATGTCTAGACATTCAATTGAAATTTTCTAGGCCATCAAAGAAGCAGTAA
AATATGACCTATAAGGCCGGGCACATTGGCTCATGCCTGTAATCCAGCA
CTCTGGGAGGCCAAGGTGGGTGGCTCACCCGGAGGTCAGGAGTTCAAGAC
CAGCCTGGTCAACATGGTGAGACCTCATCTCTACTAAAAATATAAAAAAT
AGCCCAGCATGGTGGTGGGCGCCTGTAATCCAGCTACTCAGGAGGTTGA
GGCAGGAGAATCGCTTGAACCTGGGAGAAGGAGACCGCAGTGAGCCAAGA
TGGCACCATGCACTGCAGCCTCATTAGAGAACATCGGGAAG
>Contig42
GAAACTAAAGGCTTATTTAAAGCGCGAGACCGTGGCGCCTTTGGACTGGA
CCCTTTCTAATGATCATTAGTATCAGGCTATGTGGGAGTTGACCGTTTT
GCATAGCCTGAAAGCCAACAGTATCACTCCTCCTTAGGTGTGGCAGAGA
TGTGAGAGAAGGAGACTGACAGTCTGTGGGTGTGTATGCAGTGTGGGG
AAGCGAGGCACAGGGGACAATACTGTGGTGTAGAAAACCTAGTCTAAGGTA
GCATCAGGAAATTCATGAAACCAAATGAATTCATAACAGCACAAGACA
TTATTTGTTTTTGCCTCCCTCTCATTTTTTTTTTTTTTTTGAACAGAGTC
TTGCTCTGTATCCATGCTCGTGTGCAGTGGTGAATCTCGGCTCACTGC
AACCTCCACCTCCAGGGTTCAAGCAATTCTCATGCCTCAGCCTCCTGAGT
AGCTGATTACAGGTCTGCACCACCCCGCGCTAGTTTTTGTATTTTGTAG
TAGAGATGGGGTTTTGTAAATGTTGGCCAGGCTGCCCTGTCATTTTTTTT
TACTAGTGTCCAGTGGAGTTTTTAGGGGTACATAACATGATACTGTCA
TTAATCTAATGGCTAATGAAAGGGATATGTATATGTTTTTGTGTTTAAAA
CAAACCTCTTTGGGGTCTCAATAATTTTAAAGAGTATAAAGGGTCTTG
AGATCAAAGAGTTTGAGTTCTGCTGGACTGGGACAGTGGTTGTCAACCCA
GATTGTACATTAGGGTCACTGCGGAAGCTTTAAAAATAGTACTGATGCCCA
ACCTTACCGCAAACCAATTAAGCCAGAATCTCTGTGGATGAGAAGTCTTC
ATTGTCATCATCACCATGACCATCATCATTGTCAACCGTCACTACACCATT
ATCATCATCATCATATCATCTTCATTATCATTGTTAGTATCTCCATCACC
ATCATCAGCATCACCATTATTATCATCATCATCATCCCCACCATCATCCT
CATCGGAACCTCACCTGCATGGAGGACAATCCACTATGCATTAGGTGCTA
TGCTATTTGCTATACTCCTTATTCTCAAACTGCCAGAGAGGCTGATAT
TATCTCACTTTATAACAGGAGGAATCTGGATCGGAAAAGTTAAGGTAAGC
TAATTCACAGAGCGAGAAGAGATAGAGCCAGGATTGAAACAGTTCTCT
GCTACATCAATGTTCCAGTCTTGCACTATTGAGAACCTCTTTAGTTAT
GCTTTCACCCCTCCAACACCACAGTAAATTTTTTTTTTTTTAAAAAAT
TATACTTTAAGTTATAGGGTATATGTGCATAATGTGCAGGTTTGTACAT
ATGTATACATGTGCCATGTTGGTGTGCTGCACTCATTAACTCGTCATTTA
CATTAGGTATATCTTCTAATGCTATCCCTCCCGCTCTCCCCACCCCATG
ACAGGCCCTGGTGTGTGATGTTCCCCACCCTGTGTCCAAGTGTCTCAT
GTTCACTTCCCACCTATGAGTGAGAACATGTGGTGTGTTGGTTTTCTGTCC
TTGTGATAGTTTGCTCAGAATGATGGTTTCCAGCTTCATCCACGTCCCTA
CAAAGGATATGAACTCATCCTTTTTTATGGCTGCATAGTATCCATGGTG
TATGTGTGCCACATTTTCTTAATCCAGTCTATCATTGCTGGACATTTGGG
TTGGTTCCAAGTCTTTGCTATTGTGAATAGTGCCACAGTGAACATTCATG
TGCATGTGTCTTTATAGCAGCATGATTTATAATCCTTTGGGTATATACCC
AGTAATGGGATGGCTGGGTCAAATGGTATTTCTAGTCTAGATCCTTGAG
GAATTGCCACACTGTCTACCACAATGGTTGAATTAGTTTATAGCCCCACC
AACAGTGTAAGCAATTCCTATTTCTCCACATCCTCTCCAGCACCTGTTG
TTTCGTGACTTTTTAGTGATTGCCATTCTAACTGGCACCACAGTAAATTT
TTATAGATTTTATAAGCAAATTTGATTTTACTGTGCAAGAATTGGTTTATT
TTTTAAACCATGTGTTGCAAACATACAATGGTTAATTGTGATTTTGCTC
AGTACAAGATCATCAGATCACTACACAGACTTGAGGTAATTCACCTAAA

FIG. 4 (20 of 61)

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AGCAAAGAGAACTGACCCACATTAAGTGAAGAGTCTTTACTTATTTA
CCCTATAAACGAGCCAAATATGAAGAGAAGGCCTTAATGTGGTTAACTATG
TAATTTTTCTGACTTTTTGAAATACTGAGAAGAGCTCATGACTCTCCC
ATCTCCTAATTCTACCTTGGTGGATTTAGACTGACCACAACCTCATGGGT
AAATGAGGGAAGACGAATAAGAAACCTTGCTTTTTTTTCTCCTTGTTTT
TGGCTGGCTGCAGTGGCTCACACCTGTAATCTCATCACTTTGGGAGGCCA
AGGTGGGAAGATCACTTGAGCTCAGGATTTCAAACCTGGCCTGGGCAACA
TAGTGAGACCCCATCTCTAAAAAAGGCGACGG
GCGGTGCGTGCTGTAATCCTACCTACTCAAAAAGCCGAGGTGGAAAGAT
CACTTGAGCATGGGAGGTCAAAGCTGCAGTGAACCTTGATTGCACCACTT
CATTCAGCCTGGGTGACAAAGCAGGACGCTGCCTCAAAAAACAAAAAC
AAAACCTTAATTTTTGGCTATTCTTTCTGGTAAGAATGGTATAGAGAT
GGGATGAGGATGGCTATTGTATGAGAGAGCAAACAGGGTCCAAGCAGTG
CTCTGGGCTGTCTAAGGACCAGTAGTCAGCTTAACTTCTCAAATTTCCAG
GGAAGGAGTTCGGAGTGGTAGAATATCCTGGGTATGCCAAAGCATCACC
TTGCAAAATAGCCTGTCTGAATAATTTGTTTCATTTGTTATGACTGGAAA
CTGGCTTTGTGTATGCCAGAGAATGGGGGCAGGAAAGAGAGATTGGTGTC
TTGAGCTCTCTGTGCCTCTGGGGCAGTGATGCTTTTCTCTCATGTGGAA
GGAGAGCATGACTGAAAAGGTGCACAAATAAGGTGTCTGTGAGAGAAAT
AACCTTCAGATACAGAGACACAACCTTCCCCAAGAGGTCTCTATTGCTC
TGCTTTTTTCTTTTTTCTTTGCTTGTCTACCATTAATAACAGAACTGA
TTATGACCTCAAAGAGAGGAGAAAGCGACTCTCCCACTTAGAGCTAG
TTAACCCATATCTTCTTAGATCTCAGTTCAAGAGTCACTTCCATCCCC
AATAAAGCCCTTGAGTGCTGAGCACCTCTCCGTATAGCATTTGTCTTA
GGGTTTTTGTACATTTCTTGTGTGAAACTTGGGTTGACATCTGTATTT
CCGACTAGATTACAGTTTCTCAAGGTAGGGATGTCTTGTGCTTGCATTT
TCAGTTCCAGCATCTAGACAGTACCTCAAGCAAACAAGGCCGAGGGGGT
GCGGATCACGAGGTGAGGAGTTGAGACCAGCCTGATGAACATGGTGAAA
CCCCGTCTCTACTAAAAATATAAAATTAGCCAGGCGTGGTGGCAGGTGC
CTGTAATTCAGCTACTCAGGAGTCTGAGGTAGGAGAATCGCTTGAACCC
GGGAGGTGGAGGTTGCAGTGACCTGAGATCCACTGCACTCCAGCTTGGGT
GACAGAGCAAGACTTCGTCTCAAAAAAAGAAAGAGAGAAA
AGAACATCAATGAATGAATGAGTGAGATGAATGAGTTAGCAGTGTTGGA
TTTAAGTGTCAGATTCTTCCAGCTTGACTTTTTCTTTGGCTTAGTGAT
TTTGAGGTCNCAAGATTTATTTCTTTTCAAAAGGTGATCACTACCATA
AGATCTTCAGAAAAAAGATGTGGCAAGCCANGTCTCACTAATGCAATCT
CTATACAACTGTATCAGTACT

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GAGGTGTCTATAAATATGGACCGATAGATGAATACAGGTAGGATGGGACAC
AATCTAAGATCCCAGGGGGGGGAGACCACACGCTTGGTTAGGGAGACCCA
AAGTGGACCGTGTGGCCAGAAGAGTCCCGCACTGCACTCTAGTGACAGTG
CAGAAAGTCACTGTGGGAAATCTAGAAGTTTCTACAGGTTGCTATTTTAT
CATAGCACTGTGCAGGCCAACCTTCTGCTCCACTGGCTGTGGGAAAA
GCTTTCTCTTTTCTTCTAGCCAGGGAGCTCTCAAAGTGTTCACCTCTCT
CACCTCCACCCAGGCGTCCAGGTGTGGAGGACACTGCCGCTGCTTGTCT
TGCTGACTCATCCCTTGGTTTTCACTTGGAAAACCTACCACAGCTGGCCT
CTTTCCAAGCATCAGCCTCCTCATTTTCTTAATCCCTTAGGTGTGATCTC
ACCTCCACACAGTAGATTGCCTCAAGGCCCAATTCCAATATGAATAAAAA
TGATTATTTTGTATCTTCCAATCTTCTTTTAAAAATATTATTTTATAAT
TCCCTTTAGGAGGATCACCTAAGTGAAGACTATTTTACCTAAGAAATGT
TAAATGTAAAGACATGGTTGTAATCTGGGGATTCTGTGTTAAATGGCTA
GCAGACAGAAGTCAGACGACAGGCTAGAAATGTGTGAAGAGTGGTTGCCT
TTGAAAGGCGGAGTTGGTAATGATTTTCTTCCATTTTCCATGCTTTCCA
ATTCTCTACAAAGGCCTTAATATTACTTCGATAACCAGGACCTCTGATAA
CCTGCCCCCACCAGTAAAGACTTAGCTGGGAAAGTCAGCTTCATGTGAG
GTAAAAGGAACAGGTAATACACAATTCCTCACTGCCAACTGTCCGGTGTG
CAGGCTTCACTTCTGTCATGTGGGAGGAAAGAGAAAGAGAGAGAACT
CCAAGATCCAAGATCCAGCAAGAAGGCTGGAGTCTGAGGACGCAGAAA
GCTGAATGGCACAGTTACCACTATTGTGCTGAGGTTCTGTGGCCTCTGGG
TCTCTTGACAACCTGGGCAAAGACCCACAGAAAATCTCTAGACCCTAC

FIG. 4 (21 of 61)

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CTGTGGGAGGGGAAAGTCTTCAGATCACTACAGGACAGCCACCTGGAA
CTCAAATGGCTTACAGTTCCCTTCATCCAGAGGGTCTTCATCTAGTACATA
CCAGGTGCTAAGCCTGGGTGCTGGAGACATGACGGGGAACCCATTTACCA
TGGCTTTGTTACTGTGACATTACATCTAGGGAAAGCCAGCAAAGGGGAG
GGATCGAGGAGAGCTTGTAGGCAGAGAAAATACCCAAGGGCAAGGGAGA
AGCCAGCCTGTTCTGAGCACACACAGTGGTTCATCTAACTGGGCCTCAG
TGCCAGGTTGGACTGGAGATGGGGCTGAGGAGCTGTACAGAGCATTCTG
GACACAGATGTACATAGTCCCTTGAGGTTAGGGTCTTAGGCATGGCAG
CATTGCTTTGAGTTTTCTTTTGTAAATGTTGCCATTCATGACAATGTGG
AAGATGGGTCTTGCAGAGAAGGGCAGGGCTGTGAGACCAGTTAGGAGAC
TAAGATGTGAGCCAAGGAAAATGAGGAACACCTGAACACTGGGGCAGGTG
CAGGGCCAGAGAGAAGCAGATGGCTTCTGAGGTTTTAAGTAGGTAGAA
TCAAGGCAGTGGTAAAGATCTTTATTACATATAAACTGGAATAAGCCA
TCTGCTCCAAGACAAAAGAGTAGGCGGAAAACAATACAAGACAGAAATGG
AATTAGAACAAACCTGGGAGGAATGTGGAATTAGAGTAGAGAGTCCAACA
CTGGCTGCAATCATAAAAAATGTAACAAACAAAAATTTGCTAGGTGTGC
TTACTTAGAAATAATTAGCTGTCTATTAAGTTCACTTGTGTTATGGCTT
AAATGTGTCCCCAAAATGTGATGTGTTGGAACTTGATCCCCAATGCAA
CAGAGTTGAGAGATGGGACCTTTAAAAGGTGATTAGGTATAAGGGTTCT
GCCCTCATAAATGAATTAATACTGTTATCATGAGAGTAGATTCTTGATAA
AAGGATGATCTCTGCCTCCTCCCCACAGCCCTTGTGTCATGCTTCTCTG
CCTTTCCACCTTCTGCTATGGGATGACACAGCAAGAAGGCCCTCACCAGA
TGCAGCTCCTTGATCTTGGACTTTCAGCCTCCAGAAGTGAAGCCAAAC
AAATTTCTGTTTATTATAAATTACCCAGTCTCAGGTATTCTGTTCTAGAA
GCACAAAATGGACTAAGATCATTAGATTATCATTTTTATCAGACTGTTG
AAGTGAAAAATAAAATCAAATAAAGAAATTAAGAGAGCTGCATGCAGCA
GCTCATGCCTATAATCCCAGCACTTTGGGAGGCCAAGGCAGGTGGATTGC
CTGAGCTCAGGAGTTTCAGACCAGCCTGGGCAACCGGTGAAACCTGTT
TCTACTAAAATACAAAAAACTAGGCCGGGCGCGGTGGCTCACGTCTGTAA
TCCCAGCACTTTGGGAGGCCGAGGCGGGTGGATCATGAGGTCAGGAGATC
GAGACCATCCTGGCTAACAAGGTGAAACCCGCTCTCTACTAAAAATACAA
AAAAAATTAGCCGGGCGCGGTGGCGGGCGCCTGTAGTCCCAGCTACTCGG
GAGGCTGAGGCAGGAGAATGGCGTGAAACCCGGGAAGCGGAGCTTGCACT
GAGCCGAGATTGCGCCACTGCAGTCCGCAGTCCCGCCTGGGCGACAGAGC
GAGACTCCGTCTCAAAAAAAAAAAAAAAAACTAGCCAGGCATGGTGGTGT
GTGCCTATAGTCCCAGTACTTTGGGAGGCTGAGGCAGGAGAATTGCTTGA
ACCCAGGAGGTGGAGGTTGCAGTGAGCTGAGATCATACCACTGCACTCCA
ATCCAGCCTGGGTGACAAAGCAAGACTACATTTCAAAAAAAAAAAGAAAG
AAAAAGAAAAAAGAAAAAGAAAAAGAAATTAAGAGAAGGGCAGGTATTAA
CCCCAAATATCCCACCATAGGGACACATTAAAGTTTGCTTGGCCACTCCC
CTAGCATAATATATGGAATGTCTTCAAGGACCCTCTGTTGTAATACAAG
GCCCTGCTGGACTTAATACAACCTGCAGGCTTTGAGATCCCTACTCTGTT
GCCATCTCTCATAGGATTTGCAGACCAATCCAAATACTTAAATAGCAA
CACTCACAAACATGCAAAATCAGAGCAGAAAAGAACTTCTAAAAGGCCCT
GAAACTACACTTTATGAGAGAAGACAATAGGGACCTGAGGGTGGTAGAAT
TTTCTCTCTATGCATCTATGTTTCCAGGGCTCACTTTCTCAATAAACTCT
TAAATTGCTTTTAAAGTAAGGGAACAAGCAAAACATTACATTTAAGAGAAA
TCAATTTTATAAAGAAGGGGGGATGTCCAGGGTACTTTGCTTCCATGTTT
TGCTTCCATGAATTTGTGTTTAAACAGAAGATGCAGAAAAACACACAATTA
TTGCAAAATCAAGGAAATCCACTCTAAACATCCCTTGTTTTCCAGGCCA
GTGTCACAACTGAAAACACATATTGTGGCTAATTATGTGTACAAATTAG
AATGACAAGGCAAGAAAAAAACTCTCTGATTAATAATAGCAGCCAA
CACAGACAGCCTGTGTAGCTCGACTCTGCTGGTTTTATAAAGGCAGAAGA
AGCAAACGGCTTCTGTGACCGCAACAGGAAGGGCCTCTGCTCTTAATAAA
TAAATAACATTTAAATTATTCTCCCCATTTGCAAAGCATTTTCCAACCTC
ATTATCTCATCTGACCAGGTATTATTGTATCTGACCAAGAACTTGTATAC
NAAATAAAGAATAAAAAATAAATATGGGCCANGCACAGTGGCTCATGCTT
GTAATCCANCACCTTTGGGAGGCCAGGCGGGTGGATCACTTGAGGTCAG
TAGTTTGAGACCAGTCTGGCCGACATGGCGAAACCCGCTCTACTAAAA
ATACAAAAATTAGCCCGGCATGGTGGCACATGCCTGTAATCCCACTACT

FIG. 4 (22 of 61)

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TGGGAGGCTGAGGCACGAJAATTGCTTGAACTCGAGAGGCGGAGGTTGCA
GTGAGCCGAGACTGCGGCCATTGCCCTCCAGCCTGGGCGATGAGAGCGAA
ACTTCATCGAAAAACAAAAACAAAAACAAAAACACCTTAGAAGA
AGCGTTCCTCCTCTTGTCTTCTGAAGACACTCTACGCTGAAACAGTAACT
TTCAATAAACCATCTCTTCTCACCGCACTCTGCGACTTGCTTGAATTCC
TTTGTGTGCAAGATCCAATAAGCCTCTCTTGGCGTCTGGATGAGAACCCT
TTFTTTTGAATACTCTGACACAACAAATTGCAGAAAGAAAGTCTCACATG
TATAAAATAAGCAAAAAGATTCTCTGGCATCTGAAGAAACAATTTCTTG
TCAATATTAGTATCACTATAAGTGTAGAACAACCTGTTGTATGATGCTAC
ATAAAGTATATGAATCTGAATACTGTTGGATACAAAGGGAGACTATNNAA
TGTAATACGTGCGCCGAAATGACTACACTGTTGGTGATCTTTCTTTCAAG
AAGCANAATATTGCCTCNAACATCCTGTACATGGTATAAAATTTTA

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CCCAGCAAGAACACCAATACAACGGGGGGGGCGTTCTTTGTGAGGGGTGG
GGAGGTCAATTTTTTGGAACTGCAGCAGGTAACACACAAAACTTCCACA
GCTGCTACCAGCTTTCCAGGAGAGCCTGTGTACCTGGAGAGGGAAGGCA
AGTGCTTCCGAACCTTGACTTGATGTCTTAGATTCTGCAATGCGTAGTCTG
TAGGGACAGGCTGTAGCTTATCCTATAGGCTTGGGCTGGAGTCAGCAAGC
ATCTGGGCTGGCAGAAGATAAAAGATGCAAAGGTGGAGGAAAGCATACGT
GGTCTGGAAGACAGACTTGGTGGGTGGGTGGCTGCTACAACACCCCTAGTT
AGAGGTAGAGGGGTAAGTCAGTGTGTCTTCTGCACAGGCCTCTTCCCCAC
CTCATTCTTCATTTCCCATACAGCCTTGCTGAGTTATTCAAAACATCTG
ATTCAACTGGAAGCTGGGTGAGGATGACCTAAAGGACTAGTGTGATGCC
TGCCAGGGGTGTGGGCCCATAGTCAGAGTCCAGAGCCTCCTCTCAGCTT
TTAGCACATCTCACCCACATCCTGGGTCTTAATTAGCAATATGAAAGCA
AGCCAAGTGACAAGATTTTGTCCCTGGGAAGTCCAGAAGCACTCCTTTTC
TCATTTGTATAAGCATAATGATTTGCTTACATAAATAATCATGAAAATTC
AAATCCCTCTCAGAAATCAGGTCTATAAAACCATGAAATGCAGCATGTGGG
CAAGAATCACAGGGAAGGTAGGTCTTGGAAAAGAAAGGATGGCAGGGAG
GAAGAAAGCAGGGTGCCAGGGGCCCTGGGCTGCTGTCCAAGTCAGGTGGC
TCACCGTCTCTGAGAACATTTCACTTTCTGGTAAATGGGGCAGTTGGAGA
TAGAAGGGTTGGGTGAATGCCAAGAGTGAGCACAGCTGAGGTGAGTGTG
TGCCCTGCAGTCCAGGCGGGAGTAGAAATCCTGGGCCCATCTTACCTCCGA
CCTCATTTCCTCCTCTGTAATAATGTGGGGGTGGGGGAAAGTTCTGGTCA
TCAGCCCTAGCAITCCATGGTTTCAATTCCTCATCAGTGTGGAATCAC
CAAGCAAGAGAAACAGGATGGAGAATAACCGGATGGGTGCAATCGGAGGTG
CTATTTCAAGGTGAGGTGGCCAGGGAAGGCCCTCTGAAAGGGTGGCTTGAG
CAGGTGGCTGAATGTACAGAAGCTGCCAATCATGAAAGATCTGGGGTACA
GCATGCCAAGCAGAGGAAATGCGAGTGCAAAGGCCCGAGATTGGATGTG
GGCTTAGCACAAATGTGGCATGGCAAGAAGGCCAGTGTGGCTGAAGCAGC
ATGAACAATGGGTGGAGGGGCTGAGAGGACAGAGGAGCAGGAAAGAGCCA
GGCTTGGGTAGGAGAGGTGTCAACTTGATATATGATGCAAAGCCCTTGGGA
GGTTCCCAACACAAAGCAATGATCTAATATATGGTTTTAAAAATGCCA
CTCTTGGCCGGGCGCGGTGGCTCACGCCTGTAATCCCAGCACTTTGGGAG
GCCGAGGCGGGTGGATCATGAGGTGAGGAGATCGAGACCATCCTGGCTAA
CAAGGTGAAACCCCGTCTCTACTAAAAATACAAAAATTAGCCGGGCGCG
GTGGCGGGCGCCTGTAGTCCAGCTACTCGGGAGGCTGAGGCAGGAGAAT
GGCGTGAACCCGGGAGGCGGAGCTTGCAGTGAGCCGAGATTGCGCCACTG
CAGTCCGCACTCCGGCCTGGGCGACAGAGCGAGACTCCGTCTCAAAAAA
AAAAAAGAGTGAATGTCAAAAGTGAAGCAGACCACTCAGGAGGTGAGG
GCAATGGACTGTGCAGGAGAGACTGACATCTTAGACTCGGGCAATAGGAG
AGAAGGTGGTGAGGATTATTTCTGGGCATAAAGGCAACAGAACTAGCTG
ATGGCGTCAACGTAGGAGATGAGGGAAGAAAGAAATCAAAGGGCATTCA
TAGGTTTGAAGGTGAGTAACTGGGGATATTTAACAGAAATGGAGAAGTC
TGGGGAAGGGGCAAGTATTGTGGGGGCGGGGTCAAAAGTTCTGTATTTT
GGCCAAGTTAATTAATTTGAGATACCTCTTAGGTGTCCAAGTGAAGAT
GTCAAACAGTCAATTTGAATACAAATCTGAATCTTAGCCAGGATGGTCT
CACACCTGTAATTCGACACTTTGGGAGGCTGAGGTGAGAGGATCACTTG
AGGCCAGGAGTTTGTGATCAGCCTGGGCAATAGAGCAAGACCTGTCTCC

ACACACACACACACA. AAAAAGTCA. CAGGCATGGTGGCACATGC.
GTAGTCCCAGCTACTCAGGAAGCTGAGGCAGGAGGATCACTTGAGCCCAT
GGTTCAAGGCTGCAGTGAGCTATAATCACATCACTCAATACTACACTCCA
GCCTGGATGACAGAGAGAGACCTCATTATTAAAAATAAAATTTAAAAAA
TTAATTAATAAAATAAATCCAAATCTTCTGAGATTCAATTAGGAGTAA
CTGTCTAGTAGAAGGCATATAATGCCATGGGTACATGATACCATCTAAT
GAATGCCACTGGAAAAGAGAGAATAGCTAAAAACTGAGCACTGGGCACAC
CAGCACAGTGAAGGTTGGAAGGAAGAAATGGAGCTAACAAAGGAGACAAAA
GAGGAGTAGCCAGTGAGAAGAGAGAAACATCTGGAGAGAAGAGAGAGCAG
CAAAAGGTGGGTGAAGGAGAATGTGGTCCACCAGGCCCAACAATGCTGAG
CAGTTGAGTAAGTGAGGACCTGGCCACTGAATTTGGCAAGAAAGAGGATG
TCAGCGGCCCTAGAACAAAAGTGAAGAAGAGCTTGAGGACGGAAGCCTGA
CAGGAGTGAAGTGAGGAGAGAATGAAAGGTGGAGACATGGAGCCAAGGAG
CACTGAGACTCCCTTGAGTAGTTTGTCTGTAATAAAAGTGAGTGCAGA
GACGGGGCAGGGGGACAGAGAAATGCAGGGGTAGCTGGAGGGAGCCACAG
AATCAAAAGAGGGTTTGTGTGTTAAGATGGTAGTTGTACATAGCACAT
TAGTAAGTTCATGTGAATCACACGTAGGTGAGACAGATCACTAATGCAG
GAGTCAAATCCTTGACAGAGCCCCAGAGGAGGTGATGAAGGGAAGTGATG
GACATCATTGAGATGCAAGTAGGTTAGCAATTCCTGGGGTACAAATAGGA
GGTGACTCCTTCTGATTGCTCCTGTTTTCTGAATGAGATAGCACATAAA
GTCCACTCAGCCATGTTAGCTGTTGAAGTCTTGTGGCTGTCTGCTGT
ACAGACTGGGCTCTCCTCTCCAGCATTTCTCTCAGACTAAGCTGAGCTG
CACTAGCCGCTGCCACATCCTCTTGGGGCCATCCTCTGCCACACTCCACA
TATTGCTGTGGTTTGTCTGCAACCCCTGGAAGGTCTACTGGCTGCTCCT
AGAAGAGTCTGGGCGGCATCTCTCCCTTACTCGTTATCACATGGTGCTGT
AAGCAGTGGCCACACACTTTAGCTGGTGGGATGGGCCATCACAGGCAGTA
AATGCGAAAGACTGCTCAGATTTTAAAGCACCCATGAATCAGTAGAATGA
GTTTAGAATTGTAGTCATCAACACACATTAAAAAAGAGAGAGAGAGAGAG
TAAAAAATTAGTTGAGTAGGATAAAGCCATAAAAGATATTAACTACAAC
CCAGATAGGAGGTGCAAAATTGTCTTACATAAATCAGATGGAAAAAGTT
GAAAGCAGATAAGATAAAATAGGTAAGCATGACATTTAAAGGTATTTCAT
GGGACGTGGTTACAAAACCAACTCACAACTAAAAAGTCTTAGGACCTCTC
GCTGACTTAGGAGCCTGATCCCACTTTGAGAATGACTCAGTGTGTTACC
CTGTGGCTAGTGTAGACCAATGATCCTGTCTCAGAGTCACTAGCCAACAG
CCCATATCAAGTAATTGAACTTTGACTCAGAACTCAGTGTGAGAACC
TTTGACTTAGGAACCACTGTAGTGGTTAACTGCAATTTGCACCCCTTAG
TTCAGGGCTTTACAAACACCGGGGGAGGGGGAAGGCATAGAGCTGA
TGACCTAAAGGAAACCCATTGCAGCAACGCTTTTGTGTTAAGTTTACAAA
TAAGTGTGTTTGTAGAACTCTCCAGGTAATGCCTTTGTTATTTAATGTGT
CTGAGACAATTCTGCACATTAAAGAATATAAAATATTACCTTGTAATTCC
AATTTGAAATGTGTAATTGACATTAGACTTCTATTTTAAATTTGAAATGTC
TAAACCAATGTGGTTAAGTTTGTAAAAGGTGTGTGAATTTTGTGCTGAT
TTACTACATTTTTTTTTTAAATTTCTTTTTTTTGGAGTTTATGGGATTGC
TTAGATGGCTAGAAAGATCGCTAGGCACATGTCC

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GATGTGTGTACGTGTGTGCAAAATACCGTGCCTTTTTTGTCTTTTGT
GAAACAGAGTCTCACTCTGTGCGCCAGGCTAGAATGTAGTGGCGTGATGT
CAGCTCACTGCAACCTCCGCCTCCAGGTCCAGTGATCTCCCGCCTCA
GCCTCCCAAGTAAGTGGGATTACAGGCGCCCAACACGCCCAGCTAAT
TTTTGTATTTTGTAGTAGAGACGGGTTTCCCATGTTGGCCAGGCTGGTC
TCTAACTCCTGACCTCGAGATCCACCCACCTCGACCTCCCAAAGTGCTGG
GATTACAGGCATGAGCCACCATGCCTGGCCAATACTGTGCCATTTTATTA
TCAGGGACTTGAGCATCCATGGATTTTGGCATCCATAGGGGTCTGTAAAC
CAATACTGCACAAATACCAAGGGACAAGTATTCTAAAAAGACCAAAAA
TTAATAAGCAGGACGCTGAAGGTAATTGCCCAATAAAGTCATGATCCCT
TGCCCAAGTGTCTGAACCTCAGCCAGTTTTCTACTCAGGACCTATTGGCT
GCAGAGGTGGTAGGAACCATATGAGAATCCTGCAATATCATGGCAAGTAT
GCACCTTAATGATATCTGCAGTCTTCCCCAAAAGGACCTTACATTTACC
ATACTGCTATGCTGCGTGAGAGGGTAATACTCAGATTTTTTTTTTTT
TTTTTTTACACAACGCTCTTACTGTGTTGCCCACTGGAGTGCATGGCT

FIG. 4 (24 of 61)

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CGATCTTAGCTCACTGC .CTTCTGTTT .TGGGCTCAAGTGATTCTC
GCCTCAGTTTCTCTGAGTAGCTGGGATTACAGGCGCCCGCCACCATGCCTG
GCTAATTTTTGTATTTTTAGTAGAGACGGAGTTTTGCCATGTTGGCCAGG
CTGGTCTTGAACCTCTGACCTCATGTGATCCGCTGGCCTCCCAAAGTGCT
GAGATTCCAGCGTGCGCGGCCATACCCGGCCGGGAATTCTTTATATATTC
TGAAAACTAATCCTTTGTGAGACATAAGTGTTGTAAATATTGTATCCCAG
TTTGTGGCATGTATTTTTAATTTTAAATGGTGTCTCTCAATGAAAAAAGC
TTAACTTTAAATGAGGTCAAATTGATCACCTTTTTATTTATGGTTGATT
CCTTTGGTGTGCTAGTGTAAAGGAATGTTGTTCTTCTCTGTCCTCAAGTTGC
AAAGATTTCTTGTGTATTTTGTCTTAAAGTTTAAAGTTTGTCTTTTCC
CATCTGTGCACATTTACATTTGCTACATCTCACTGACTGCTTCTCTGCTGC
TGCAGAGCAAGCTCCATGAGAGCAGGAGGCATGGGTCTGCTTCTTGTG
GTCCCCAGAGCCCTATGTGATGACTAGGACCTGGCAGGGGACTAGTGAGT
AGCTCCTGACTAACTGACTCAATGAATGAATGATTGGATGATTGAACAAA
TGGGTATGGGAGTTACAGCGAGTAAGAGATGCCCTAGAAAGAGATGAAGA
AGGAGATGGTATAGGGTAGTGGTTCTCAATTCTGGGTCCATGGTGGACTC
ACCTGGGGACCCTTAAAAATGTACCGTGGAGGATCCAGCCCAAGAGATTC
TGTATGACTGGTCTAAGATGTGGTCTGGGCACCAAGTGATCCAGTGTGC
AGCCAGGCCTGAGGCCACTGGATTTGGTGGTAAATGAGGTAACTATCAAG
GGTACAGACGTTGGTTGCCAACAGGCTTGGGCTTGAATTTAAGCTTTGTC
ACTGACTTGCTGTGTCTCTCTGCACTCGTTGAGCCTGTTTTCTCAGCTGA
GAGATGGGTGTGATAACACCTACCTGCTGTAGTTGTTGTGAGAGTTAGAG
GAGATAAGCATGTTCTTGAATGAAGTGTTCTTAATCCATCATAGGTT
TTTTGCTTGTGTTTGTGTTTGTGTTTGTGTTTCTTTTCAAGAATGA
GGTTGAGCCAGACTTTGACAGCTGGGTGGGAAGTGAACATGTGGTGATTG
GGAGAGAAGGGCAGTTTATGTGAAGGGAATGTAATAATTAGAGAGTGGGC
GTGGGAAGACATGCTGGGGAGAGTGAGCAGGCCGTTAGCCCTGGTAGAG
GGTGCAAGAGAGCAGTGCGGAATCTGCCAGGGAGACAGGTGGGTGACCAG
GGTGCCAAGGGTGTGGCTTTTCCCAGGTTCCCATGGACACAGCCATCCTC
CCAGATGCCCAGCCTAGCTGTGAGTGAGCAAGAGTTCTGGATTGTCTCTC
TCACTCTGTCTTTTTCTCTCATTCCAGAAACAAAGCAGTGACTGGTACTT
AGGAGGCAAGTCAAGTTGGGAGAACTTGCTTCTGCTCAGGGGAG
CAGAAGCAAGAATGGAGGCCCAACCATGCTGGAAGATGATGAGGGTTT
GGTTCAGGGAGGAGGAATATTGGGGATCTAAAGGGGCTGGGAGTGGGGC
AGGACCCTGCCTTAGGACAGGTAGAAACATTTCTATAAAAAATGGGGTG
GAGGTTGATGGTAGGACCAGGCATCTTAGTTGGCTCCCTGGAGTGTCAG
GCCCTTGAGATGGTCTTTAAAGCCATGCAGTGGGGTTTGAATCTGGTGT
TCAAGCTCATAGGTTATTAAACATAATGACACTTGAAACTATTGGGAGA
GCTCAAGTGAGTGGCCTGGAAGTTCTGTGTTGGTGACGAGGTGACTTAG
GATGTGCTGCTCCAGACTCATATCTTGACTGCACACCTGATGCTTCATC
TGGCTATCCTGTAAGCACCTTCAACTTAACATGTCTACACAGAAGCTCTT
GATATTCCTGTTCTCTCCCCAGTTCCTCAGTTCCTTACCAAATGTTCTTCC
AGTTACCCAATTGCTCAAGTAAAAATCTAAGTCTTCTCTTGGATTCT
GCCTGTTCCCTCAACATCCCACCTATCCATGAGTGTCTGTGGGCCCTGC
CTCTGAAATAAATCCTGCCTTTGTCTCCAGTTCCTCCAGCCACCCATC
CTGGGGCTGCACCCTCCTCCTTCCAAGCCCTCTCCCTTTCTTCTGGTG
CTGCCTGTGATGTCAAGCATATGCATCAGTGGCAGGACATTTGAAAT
GCAACCAGTACAATTGGGCGCGGTTATGCCCTACCAGTTTTCTTCTTAA
ACATTTTATATTTATGTTTGAAGCATGCCACCTTTCTTCACTTGCCAAC
TTGACAGATTTATTAGTTGACAACATCCGCTGATAGCATCAGTAATAAGT
TAATTGTTTTTGCACATGTAGCTTTAATTATTCTCATTATCATTATAGG
AGTTATTCTTTGTAAAGGGTAACTGAGTTTTCCAAAACAAACAGAAATTT
GGGGTGGGCCCATGGAGCGTGACTCATGAAATCAGATTCTTAGAAGGACC
TCGGCAAGTCTCTGGGTTGCTGTTAATGAGCCTGGCTGGCTGCCAGGGT
GTGTCTGCCCTTTATGAGGCCACCACTGTTCAAATGCTTGCCTGCAGCAT
TACTTGCTAGGTAGTGCTTGTGTTCTACTGAACTGTGAGGATCCAATTC
TTTGTGGTCTAAGTAACAATACTCAGATTCACAAGGAATTGATTAATAAG
CCAGAATGCCAATGTATTACATTTTGTATGAAGACCATATTTACAGTGAT
TGTATCTGCTCAAGCTCAAATTAGGATTAGAGTTCTGACAAATACATATG
TGAGAAGTATGAGGTAAATACTTGAAATTTGGACTTTTCTAGAAAATCT

FIG. 4 (25 of 61)

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GAATGTGATTGCCATTACATACCTTTCTGGGGATGATGATTCTTGTACT
TTTATTTTAAAAAGACATAGAAAATACTTAAGAATCAGATTGCTTGGCT
GGGCACAGTGGCTCATGCCCTGTAATGCCAGCACTTTGGGAGGCCAAGGTG
AGTGGATTGCTTGAGCTCAGGAGTTTGAGATCAGCCTGGGCAACATGGTG
AAATCCCCTCTCTACCAAAAATACAAAAAACAACCAAAA
AGAATAAATTAGCTAGGTGTGATGGTGCCTGCTTGTAGTTCCAGCTACTT
GGGAGGATGAGGTGGAAGAATTGCTTGAGCCAGGAGGTGGAGGTTTCAG
TGAGCTGGGGTTGCAACAGTGTACTCCAGCCTGGGCGATAGAGTGAGACT
CCGTCTCAAAAAAATAATCAGATTGCTTTATTGCTGGTTTTCTTTCT
AAAACAGATTGGGTCCCATCATCCCCTGGCCCCCATTGGTTAATGGTT
CCTCCTTTGTCTATTGAATAAAATACAGATGTCTGCTTTTGGCAACATGG
TTGAATGTAGACACTGCAGGGTCTTCTGACTCAAAATGATTTAGGCTTA
GATAAACACATTTGGAATGCATTTCTGGATTAAACCAAGGAAAGGAG
ATCTCTTTAAATCCCCTTTCTGTTCCCCCTCCCTACCCCTCCAATTGG
GCTTAAGTAAGAAGGGTGGTTACCCGCTAGTAAACCCCTTCGAAGGGGG
TCTTCTCTCTAAGGGAAACCTTGTGTTTGCATTTGCTTCAATGGGCC
CTTGATTTTGTTCCTTGCTAAACGGGTGCTAAACCAGGGCCTCCTCTT

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AAGGCTTTTAGAATATTTGCACACTTTAGAAATGGAAATGTTTTTGGGG
GCGAGTTGTCTTAATATTTTCTAGCTTGTGTGACATCCTTTTGA
AAGCAGCAATTCCTGGCCTTTGTGAGAGATGGTGAATGCCTGCAGGTGTGT
GGACCAGTCTGCTCCCTTCTTCTACATGCACGGCCCCCAGCTGGGCCCA
GCAGAGTGCTGTTACAGAATAATTTCCAAGGGCTGTGTCTTAACCTTTG
GTCTTGTCCTCCCATTTGCTGTAGATTTGGCCAATTGACTTCATAAGTGCCT
CTTATGAACATAGATGTTGGCAATGGAAGTTGAGGACCAGTCAGTGGTTG
TTTTATTGAACACACAGCGTAAATCCCAACACAATGCTGACCTAAGAGAA
TTCCAGCCACTCTGATTCTCAGTCTCTTTATATCTGAAAGGGTTCTGTTT
CACTTTTTCCAGATCAAAATGTCCCTGCAGCTACTCAGCAGAGCTGTGCG
CAACTTATACGTAGAAGAGGTAACAGTCCACAAACAGAAAGGCACAGGAC
GAGAGTGGTCTGGGTGATGCTTCTGTGGGGGAAAAGGTGATGAGGGTGC
ATCTGCACACCTATGTTTCATAGGTAAGTCTGGGAGGAGGTGACCTCCCCT
TTGGTTGAGGTGCTGAGGCGTCTTGTTAGAATGGCACTATTCCATTATC
TGATGCAGTCTGTGGGAATTTTGTGGTATGGCCACCACAGGTACCATGCT
GGGAACAATGCCAGATACTGCCTGCTAAGCCACAGCATGAGTCACATGAG
CATTTGTGGGCTTTGGGAACTAAAGTTATTGAACGATAGTTATCTGAAAA
GGAATTTAGGGAAAGGGACTTTAGTCCAGCGAACAGTTTGCAAACCAGG
GGGAAGGCAGCCTTCAGCGTAAATGAAGACGTGTGTGCCCCAAATAACA
AAGGGAGAGTTTGTCTTTTAGAGAGTAAATGTCCACGCAAGGTTCCACTT
AGGCAATGAAAGATGCAAACTTGCTTAGTTCTGATTTGTTTACATTGTC
TGAATTCGGATTGGTCCGTGCAGGCTTTTCTGGGAACTCCAAATACATGT
ATGACCTCTAGTCATACATGGCAAATGGCCGCTTGGCTCTAATTTGAATT
TAGGCCCAGTTAGTCACTCAGGATTAACCTTTTTTCAGGGTTACAGCTCT
GAACAATGGACTTAGACCTGCAGGACATAATCTGTTTCTTAACCTCTGGGAC
TACCTGTGCTTTTGAAGTGTGCCAGTGAGCAGCTGTGGCTCTGGGCCCA
GACCCACAGGGCGATAAGGCACAGAGGTACGCATGGAGCAGGCTGTCTTT
GCTGAGTGATCATGAAGATACACTTACATAGAGCAGCACTTTTCTTCCA
GTCTTTGTGATTTAACTCATTAGATCCTTATAACAAGAGTCAGTCCTCTA
TTTAACCCATGAAGCACAGGTGGAGTCCAAGCTTAGTTTGTGAAGGATGA
GCCAAAAGGATTCTTCTTGTAGACCTCAAGCTCAGCTCTCTCCATGGG
CCCTGGAGTAGGTGAGAAGGCCTCTGTCTTCCAGAGCCCACTGCCAATCA
TCTACATTTTCTGTTAGCCCAATTCTAGGACATTGCTTTACCAACTGAAG
GGTGAGAATATCATAAGTTATAAAAATCAATTGAAAAACAAAAGGTAC
AGAACAGAAAAATAAAGATGAGAATCTATTAAACATAGTGATGTTACTGG
AAAAGGGGGTCTCAAACCAGACCCCAAGAGAGAGTCCTTGGATTTACAC
AGGAAAGAACTCAAGGTGAGTTGCAGGGTGCAGGTGAATTGAGAGAGTTTA
TTGAAAGCTATTCCATTACAAAGTAGAGCATCCTCAGACAGCAAGTGGAG
GAACATGCCATCATTAAATTTTCTTATATAGGAATCTTGTCTATATAAA
GACTAAACTAAGCTGTGGCTATGTGTGGGTGGGCCGACAGCATGAAAAACA
TTTATTCTCCTATTGATTTAAAGAGAACTATCCTTGACATTTTAGTGTGT

FIG. 4 (26 of 61)

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TTAAGTACATCAAAGCAAACTATAATTACTTGAAAGCATATATTTTAA
TAGGGATTGGGACATCTGGGCTTTCTGTTGTTGTAGAAGTTTGTCTTGC
AGGGATTACCAAGCCACTTCTTAGCTGTAAACATCTTAGGGCCATGGGT
CCTGACTGGCAAGGAATGTGTCTTGCTAGTTTAAAGATGGGCTTGATTG
AAAATGGTGTCCATCTGGCTCTCCTAGGCTCCTGCTTCTTAACAGTAAG
GGTAAATGCTATGTTATGAAATGTCATTTCTGCCTTTAGCTTGCAAACCT
TTGATGGTGAAATTTCTCCTGTCCGTTTTAGTGGGGTATTTATTCTGCAT
CCACGTCTTCACAAGGAGCTGAAAACAAATTTGGATGGAAGCAACTGGGTT
TTATGGGACACGTTAATGTTTTAATGTCATTTGGTGTGGAATTCAGATGT
CCAAGCAACATTTTACACTACAAATCTGCAACTTTAATAATCACTCAAAG
TACCTGAACCTCAATGCTTTTACAGACAGACTTGGTATAAAGCCACCACCTC
TTTCTATTATGGCAGCCCTATCCTGAGGACACAAATTTCTGCAGGGCTTC
TGGCATATCTCTGATTAAACAAATGTCAACAAGGTTAAAACAAATGTCAT
CTCTGATTTGTTTGTTTTAAAGCCTGGATTTACTCATTGAATATTTCACT
CCTACTAGCATGTCTTGTAGTAGTTTTCTTCAGGGACCCTAATTATTGCT
ATTAATAATATGTGTGCAGCTACATGTTTTTTTTTTATCAATTTGCAATG
AAAACCTTTAATTGAATCTATTAGTGTTATTATTGAAAGTGAAATCT
TTTCCTTTTGTCTTTCTTCTTCTCACACATAGTGCAGACAGTTTCCACAG
GGCTCATAAAGGAATGATTCTGCCTTGTGTGAACCTTTTGCCTTTATTG
TTAATTGCACCATTTTGTGACTGGCTTCTTGACCCTGTTGTAACCAAGCT
CATAATGTACATTATTTCTTATTTTGCAGTTGTAGACACTTGAGGAAGTT
CCCATTCTTTGTTTCTTCTTCTTGTTCCTGTGATAACTTTTTCATG
CAGACATTTTTTTTTTTTTTTTTTTTGGAGCCGAGTCTTGCTCTGTCTC
CAGGCTGGAGTGCAGTGGCATGATCTTGGCTCACTGCAACCTCTGCCTCC
CAGGTTCAAGAGATTCTCCTGCTTCAAGCCTTTCTAGTAGCTAGGATTGCA
GGCGTGCACCTACCACACCCAGCTAAATTTTTCAAATTAGCCACCCACCT
GGCTAATTTTTGTATTTTTTAGTAGAGACAGGGTTTCAACCATGTTGGCCA
GGCTGGTCTCGACCAGGTGATCCACCCGCTTAGCCTCGCATAGTTGCAG
GTGCTATTCTGAGCTCAGGGCTCTGGCAGCTACAAGCCCAAGATGCGGTC
TCCAACATGTGGCCATTCAATGTCTATGGCGCCCTCTACTGGTCTGGGAA
CGCAGCTCTGCCAGTAGCTCCAGCAGGGCACAGCTGTTAAGTCGTGATG
TTCTACAGTGACCAAAGGCAATCTCTGGACTCCTTAGCCGCTAGGTCC
TCTCTGTAGCAGGACCCAGGAGAAGGCAGGGGCTGAGGATGGCTCTCTTA
GACATTTGTGATGAACCAAACGTGTGCATTCTGAAACTTCTGTGAGCAA
GCAGGTGAGTAGAGTTGGGTTATAAAAAGTCTTAGGGTCTCACTACAGAG
ATGGACTTGCTGTGTAGATGGTGCAGAGCCGCTGAAGAGTTCTACTTGGG
GTAATGGTGTGATTGGGTTTGCCTTTTAGGAAGATTCTTGGCCAGAATG
AGCGGGGCAACCCAGAGCAGGGAGTGGCCACATGTGGGTGTGCAGTTATG
GGCCACTAATCCAGGTGATAAATGGTGTCTCTGAACCTCAGGTGGGGTG
CCACATGTCTCCATCTGCTCTGTACCTTTGAGACTGGCCTTATGGGCTGC
CTTAGTGGTCTGTTGTCTCTATCTCCTGGTTGGGCTCAGGCAATGGGAG
ATCAGAGGGAGGAAAGAGAGCTTGGTTAGAGTGCACCCGCGCCCTTCAG
GTTGGCAGTGGCCACATTTCCCTATACAGAAGGCCACAGTTTCTGTCACT
GGCCCTCCACAGCCCCAGCTTTCTCAGTGGGCCAGCCACCTCCCCATCC
CTTGCTCCTCCTCCAGAGAGGGTTGTGGATTTCCACTGTGACAGTG
CCTGGAGCTCCACCATCTCCTGCTGCTTCTCTGGACCTGCCTGCAGTTT
TATAAATAACCTTTCTTACATTACCTCTAGCATGCACCTTTTGTGTGTA
TACTCTGCCCCCTGTGAGCACATGACTCATGCCAAAGAGTTTGAATTTTT
TTCTCCAGGCAACGGGAGGTCAATTGGAGGATTTTAGACATTGAGAACAGA
TGTGTATTGTGGAAATATCTGTCTGACTGAAGTGACCAGGATGGTCCAAA
AGAGCGAGAATTTGAGGCAAGCAAACCATCAGCAGGGCCAGCAGCAGAAAT
CCAGGTCAATAACAGGGAAGCTGAGGCTCACAGGTTGGATCAGGGAATG
GGAGAGGGAAGCCAAACAAATTCATGAGCATGTCACTTGCACATATGACT
TGTTAACTATTTTTATTTTTATTTTTATGTTTTGAGACAGAGTCTCGCTC
TGTCACACAGGCCAGAGTGTAGTGGCATGATCACAGCTCTCTGCAACCTC
TGCTCCTAGGTTCAAACAATTCTCCTGCCTCAACCTTCCAGGTAGCTGG
GACTACAGGTGGCGACCACTACACCCAATAAGTTGTGTATTTTAGTAG
AGATGAGCATTACGCTGTTGCCTTAGACACGG

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AATATTGATTATTTGACCAGAAATTCATGCAGCTAACCGTGACCCCTGGC

FIG. 4 (27 of 61)

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AAAATAAAATAGTGTAT...GTACGTGCATATACATGCAAAGAAATGAGI...
GAAACTAGAAGGATGTCAATCAAATGATAACATGGTCATCTTGGGGTCGG
AGTACATTTGGGGATGAGGGGAGCTGTAAAAGCAGACTTGGACCTTTTCT
TCTACCAGTACCGTGTCAATTTGAATTTTGGAAAGAAAAAACTCAG
AAGGAGGAGAAGGAGCAGGAGGAGAAGATGGATCTTAAGTGATTTGC
CCGGGAGCACCTTGAGAAGGTGAGATTCAAGTCTAGGTCTAAGCTTTCTA
ATTCCATGAGTGGGAGTGACCCACGTCCAAGAGGAAGCTCAAAAGGAAGA
TGTTCTCCATCATCTCTTGCTCATCCTAACAGCATGCAAACCATCCA
ATGCAGCTCAGAAAACCTCCCAAATTGCCAAATTTTATTGGAAACACTTAA
TGCTGTGGTTTCCAATTTCAACTGTAAAGTAGGTATGTATGCCATTGTTA
CCATTAACTTCTCAGAAATGGAGAGAGCTCTCTTCCGCCTCCTCCCCCT
CTGCTGTGGCTTTGGTGAGACGTGCACTCAGGCTCACCTGTCTCCATGAT
CTCCAGTAAGTACACATGAGCAGAGAGGCTCAGCTCAGCTCTTCTGGT
CCCACCAGGGTTGATTCTTTGAGAATTCTAGAATGCCACATCCTAGGCCC
CCCAAAGAAATCCTGCATCTTACCCCCAGAAATATGAATCATAGCAAATT
TCAAATCAACCATCGTTTAACTACTCAGAGCTGGGCACATCCAAAAACAT
ATTTTCAGTTTTTACAACAGTGCCTGGTGCATATCGGCACTATTGTGGAA
GCAATAAATCGACACGGAGCTGAAACACAAACAAATGCCAAATGTTTT
ATAACACCTGATTTTCTTCTGTTTCTTATGCAGTTAGTTTTGTTTTG
CTTAACTCTACCTCAGACCATAGTCTGGTAAACTCACCACCCAGAAGCTC
CCTTGAAATGTGGGTATGCAGCCACTAGGTGGCAGGAGAGATTTCTGC
CTGGAGGGAGGACAGCCACTCTGTCCCCGGGTGAGGCCAGGGCCACCCTG
CTACCTGCAAAATTAGCATGGGGCTTTATGAACACAGCTTCTAATAAA
CACAGGATCTGTTTATAGAGACTCCAAACACGCCTACCTAGTGATGAA
AGACTCAACTCAGCAAGAAACCTTCATGGCAAACATCTTCAGAGATGTT
TCCAACTTAAGGTTCTGAACACAGACGCTTCCCCAGAAAGCCATTGTTT
TCAGCACCTGGGAGCCTTGCTTTGCTTTGCTTACAGACTCGCTGTTCTTA
AATCACTGCCAAGATAACATCTGTCTCTTCTTACCCTCTATTTGATA
TAAGGACTCCTCACTCTTGTTGCTTCTATTGGCTACCTCTCCACAGGGA
GAAATCGCTGATTTAAACAGCAGTCAATATCCCAAATCTGGAACAGGGAAC
AGGGAAGCATTAAAAAATTGGAGAATTTAGGCCGGGCACAGTGGCTCATG
CCTGTAATCTCAGCACTTTGGGAGGTGACGTGGATGGATCACTTAGGAG
TTCGAGACCAAGCCTGGGCAACATGGCGAAACCTCATCTCTACAAAAAAA
AAAAAAAAAAAAAAAAAAAAAAAAAACCAGAAATAGCCGGGCATGGTA
GTGCACACCTGTGAGCCCCAGCTACTCAGGAGGCTGAGGTGGCAAGACTG
CTTGAGCCCTGAGGTGAGGCTGCAGTGAGCCGAGATCACACCACTGCAC
TTCAGCCTGGGCAACAGAGTGAGACCTTGTCCAGATAAATAAATTAAAT
TAATTTAATTAGAGGATTTAAGGATTTTCCCTACAGACACCTCCTTATTT
TCTCTGGCCTTTTCTGACTACTCTCCCTAACTCCCTGCTCCTCTGGTCTC
CCAAACTACTCCAGAAAAAAAAGGGGGGGAGGGACTAAAGGAAAGCC
AGGTGACAGTGCCAGTGTGACAGATGACAAAGCATCTGCCCGAACAAACC
GTAGGTCCCTGAACCTTTCTCAAGACCTGTCTGTGGACTTACCTATGAAA
ACCAGTTTTAGCAAAAACCTCCTAAGCCAGTTTATCAAGATCCCCCTTAT
CCTCAATATCCATCTGATTGGATTCTTCATCCCCCACCATTCCCCAGTGA
TGTCACCAGGCCTTTCTTCAGCAACAGTAGTTAGTGGGTGTAGCCAGGAC
GCCCCCTCACCCCTGATATGCCCTTTTAGTAATTCTTCATCCACAGGTTT
CCACCTTGCTCCTAGGCTATACATTCCCATTTGCCCATGCTGCATTGGA
ATTGAGCCCAGTTCTATACTGAGGTCTTACTTCACCTCTCGCCATAGTCC
TGAATAAATTGGTTTTTACATTTAAAACTGTCCAGCTCTGGTTGTTCC
TTGACACAGGGTAATTTTATTCCATGTGATAGTTTGCTTACCTCAGCC
TACACCCCTCAAACCTGCAACTCTATATTCAAGAACCAGACAGCCCTTC
CAACAGATAGGAAGAGGCTGCCCTGGTGCAAAGGAAGAGGCTCTGGGAGG
AAGGAGAGAACCCGAAGGCTGCCCTCCTCTAGACTGAGCTCTGGGATG
GGTGGACGATAAAACCCAGATACGTTTAGACATCTGAGCGTGGAGAGGAC
TTTGCTTTGCTTCCACAGGGACCCCAAGGAACTGCAAGCCCTCCAGAGA
CTAAAAACAGCAGAACAGCAAGAAATGGCAGCAAAGGTCTGGGCAGAAATC
ATCCTATGTGGGCACAGACACAAACAGAGTCCCCTGTGGCCCCAGGAGAG
TTTAAAGAAGATCCAGAGGCTGTCTATTCCATATCTCAGCAGAGACAGG
CCCCTGAGCCTAAAAGCTGATCATTAGGACAAGAAGGACACGAACGTGCTC
TGCAGCGTGAACCGCTGGAACAAGGCCAATCACCAGACACCAGACCAGC

FIG. 4 (28 of 61)

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CAGACACAGCCCCCGCAGTCCCCAAGACACCACGGACCCATCGCCCCCTC
ACCAATAGCTCCAGGCTACATAGACCCCCCTCCACTTCATGGATGTCCTCA
GAGCAGAAAGGGGAGGCAGGAGTGGAAACCCTGACTTGGTTTCAGTTGAAAC
ATAAAATGACTGTACTATTGAATTGCTGAAGTTTACGTGAAAGAAAT
GAGATTTAGTTTTTGGCCACAGTGCAAAATAAGAAACGAGGCTTCAACTG
AGATTAAGGTGAGTTATAGGAAATGTACTCCCTTGAAGGACCTGTGAAG
TGTGTTTCGCTATGAGAAAATGACCAGAATCCACGTTCTTAGCTGCGGGAC
TCAGGCTGACTCCTGTTTCTGGAGCTTGCAAAAGGGCAGGGAAATCCCT
GTTTCAGGCACAGTGATTTCAATGTTTAAAGAAAACAGGTGGGCCCTGG
CAATCATGATAACATGTCATAAGTTTACATCTCTGTGAGGCAGGTAGTGT
AATCCCCATTTTGC AAAAGGAGGAAACCGAGGCTGAAAGCAGCTACATGGT
CTCTTCAATGTGGCCCAAATGTTGGAGAACAGAGCTTAACTGAATCAGCA
ATTCTATACTTAGAACTGACTCTCTCTTTATTATATCTCACTACTACCTT
GATATTTGAAATATTCAACTTTTTTCAATCAAAAAATAACAATAATTTAG
GCATAATGACTACTATGTCATTTAATTTCTTGCTGATATTTCAATATCCC
ATGCCAGGAATATTGAAAGCTCAGCTCCTTAAGAGCTGACTATGGCATCA
ACTCCCAACAACCATCCTTCCAGAAATATTTTCCCTTTCTTTTGTTATA
GAGTGGCAGCTGCCCTATATGGTGACCACTTGCCACATGTGGCTGTTGAAC
ACTTGAAATTGGCTTGTGCAAAATGCAAGTGTAAAGTGTAAACACATAACC
AAATTTCAAAGACATGGCACATAATAAAAAATGTAAATATCTCATTAAAC
AATTTTTATATTGACTGTGTAAGTAACATTTTGAATATATTGGATTAAAT
ACATGGATGATGCCCCAACCCACAGTCCCTTATCAAGTCTCTACTTCA
CATTTTTGTACTTCTGACTTAGAAATAGCACTGGCGTCTAAGAGCCTATT
AATGTCGTCAATAGGTTCTTGGGAACCAATTTTAAACAAAATGACATA
TAAGAAAACGAATAACATTGAACAAAATGACATTATTCGAGGACCTGCTG
CATGTTGTTTCACTTAAAGTCAAGTGTCCAAGAACCCTATCAGTGACATTTA
GTGAGGACTTGCTGTCTTCTGTTTACAGGAACCTGGGCAAGTTACTTA
ATTCCTCTAAGCCTGGTTTATATCCCTGCAAAGAGAGAAGGATAATAATC
ACCAAGTACTTAGTGATGTCGTAAGGAGAAAATAAAATAATAATATGAAA
TGGCTGACAGTGTCCTTGTACACAGAAAGATGTGTGATCCACAGTAGCTG
CTATTGTCTGCCTCACTTCACTAGTAATGGTCCAGGGAGGCCTTTAATGT
GCAAGGAGCAGTACATTACATGTTGGACATGGGTGAAGGGAAAGACCAG
GCTCATCTAAACACAATAGGATGCTTGTGGTGTTTTGAGGAGGAATCAAG
GACTAGTTATCCACAGCTGTAACATGCATGGATCAAAAGAGATAAGGCAC
ACAAAAGACTTTGTGCTAGCAAGCATTACAAAATGCAGAGACCAGCTG
TGGGTGGTGGTGAGTCAGACCCAGCTTCCCTCTGTGCCTGGCTGAGTGGT
TCTGGGCAAGTCACGCCATCTGTCTTGATGCCCTTCCCCATCTATAGAGA
GGGAGCACTGAGGCCCTTCCAATACTGAAGTCTTTATTTCTGCTACT
TTAGAAATATCCACATTTTGGTAAATTCAAATGATCCAATGATTCCATT
TCCTAATGTTCAAACATAGCCCCAGAAACATCTAAATGAATCAAACAAAT
AAAATATTTATTGTGTATGTTTTGATTGCTGAAACTTCTATTTTAGCAAC
ACACACACACACACAGAACCCATAAGCCTTCATCTTCTTGGATAAA
CGAGCCTTCTGTCTGGCCATTTAAGTCACGATTAAGTAAATGATTTCCA
ACTCGCCTTTTGACAGCTTCAGATGGGTCTTCTGCGTGGCAGTGGCC
CTCCTGACTTATGATTTCTGTGTGTGCGCCTGTTACCACTGCAGCTTAA
CTGAGGAAACAAGAACAAAACAGCTTCTGACCCCAAGAGACTGTTGGAGG
CAAAGGCTTCACTCCCAAGAACCTCACACGTGGGGAGCCCGAGAGCCAG
CCCTGACCTTTTCTCCAGTAATAACATAAGAAAACAACAGGCACTGGCCTT
ATTTTGGATACAAAGAGTGGTGCTTTTCTTAAATCTTCTTTAGTCAGG
GCTACCCCTTCATGGACGCCCCAACATCCATGGTTCTGCTTGAGTCCCT
GCTTCCATATTCCTGCATTTCTCACTTGAATATCCCTGGAGTACGTAA
GCAGCCAGGTTTGAAGTCTTGTGCTGTCAGGCGGGTGTGTGCATGTCCT
CTCTCTCAACAGGACACAAGCTCCCCAAATCAGACGGTATGCCTCCACGC
CCCTTCCCAAGCTTCCCAAGCACCCGAGCATGTGAGGGGAGCTGGGGC
CCAGGCCATGATGGGAAGCACTCTCTGCCTAAAGACTAGGGTGTGCGCC
CTCAACTGTGGGAATGAGCCCCAGCTCTGGTGTCTGCCTCGGTTTTTCT
CTGGACAAATCAACATGAACTCCTCACCCTCTTATCCACTTTGCATAAA
CTGAAAATAACAAACCCAGGGCTCTTCTGTACAGGAAAGGGTTTTTTT
TTATAAAATTAAACAGAGATGATTCACACACCCAGGATATAACACATGG
GCCATGAATCAAGGGCAGCATTGCTCTGGTCAGCCTGTTGTTTGGGCCCC

FIG. 4 (29 of 61)

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CTTGGCAGGGCTCTCCCCAGAATCTTCCCCCTCTTGACTCCCATCANACA
GCACTCCANCTTTGTGTACAGGCGATAAATGGGAAAGGGTAAAT
>Contig48
CATTCTTAATTAGAGAAACGCTCATTAACTAGACACCCAAATCTCTGG
GGGGGATCATTCTTACAAGCATGCCCTTCTCTTAAAGAGAGCACT
TTTTTCGCAAATAATGCTGCCATGAACATACGGGGTGATGTATCTTCGT
AATAGAATGATTTCTATTTTGGGGGTATGTACCCAGCAATAGGATTGCT
GGGTCAAATGGTATTTCTGGTTCTAGATCTTCGAGATCTTCCACACCGTC
TCCACAATGGTTGAACTAATTCACATTCTTACCAACAGTGTGAAAGCAT
TCCTATTTCTCTGCAACCTCGCCAGCACCTGTTATTTCTTGACTTTTTAA
TAATCGTCATTCTGACTAGCATGAGAGACAGTATCTCGTTGAGGATTGA
TGTGCATTTTGTAAATGATCAGTGATGTTGAGCTTTTTTTCATATGTTTT
TTGGCTGCAAGAATGTCTTCTTTTGAGAAGTGTCTGTTTATGTCCTTTGC
CCACTTTTTAATGGGGGTTTGTTTTTTCTTGTAATTTGTTTAAGCTCCT
TATAGACTCACAATAACAAAGACATGGGATCAACCTAAATGTCCATCAAT
GATATAACGGATAAAGAAAATGTGGTACATATATACCATGGAATAGTATG
CAGCCATAAAAAAGAATGGGATCATATCTTTGAAAGGACATGGATGAGC
TGGAACCATGATCTCAGCAAATATGCAAGAACAGAAAAAATTTGTTG
CATGCTCTCACTTATAAGTGGGAGCTGAACACTGAGAACACAGGGACACA
GAGAGGGGAACAACACACATTTGGGGCCTGTGAGGGTGAGGTGGGGGAG
GGAGAGCATTAGGAAAAATAGCTAATGCATGCTGGGCTTAATACCTAGGT
GATGGGTTGACAGGTGCAGCAAATCACTGTGGCAGACATTTACCTATGTA
ACAAACCTGCACATCCTGCACACGTACCCAGGACTTCAAAATAAAGAGA
GACAATACTTCTCCCTTAAGTGTCTACTGTTGCTTTGCAATAAAAAETTC
CTGCCCTTCACTTCACTCTGACTTGTCCCTGAATTCTTCTCGTGATGGT
GTCAAGAACGTGGACACTGGCTGGGGCTGGAGACTCACCAGCATCCGGAG
ACCCTCCTGAGCCCTCCAGCAATACAACTTTGACACAACTATGAAATCA
CAGATCCAAGAAGCTCAAAGAACCAAGCACAGGAAACATGATGAACTA
CATGAAGGAACATCAGAATTGAATTGTTCAAAATCAGTGATAAAGAGTAA
ATCTTAAAGCAACCAGAACAAAAATCCATCATATACGCAGAAATAAAG
ATAAGTATGACAGCAGATTTACAAATAGAAAAAAAACAAGTGCAGCAAC
AGAAACAAACTATCAATCCATAATTCTATACCTAGTGAAAATTTCTTTCA
AAACAAAGGTGAAATAAAAAAATTATTTTCAGGAATACAAAAGCGAAAAA
ATTAATCACTAGCATTCTCACTGCAAGAAATGTTAAAGGAAGTCTTTA
GGCAGAAAGAAAATGATACAAGGTGAATATTTGGATCCCTGCAAGGAAC
AAAAAGATCCAGAACTGATACTTAATGGGTAAACATGTAATTTTCATCA
ACAAGTGAATGAATAAACAAATCATGATATATCCATATGATAGACTACTA
CTTAGAATACAAAAGAAGAACTACTTATGCATGTGATAACATGAATGATA
TTCAAAATTATTATTGAGTGAAAGACACCAGATCAAAACAAAGTACATAC
TGTATGATTCTGTTTATATAAACTCTATAAATTGCATGCTCTTCTATAG
TGACAGAAAGAAGATCAGTGGCTGCCTGCAGACAGGAAGAGATTACAAAC
GGAAATGAGAATTCCTTAAGAGATGATGGACATGCTCATTACCCATCATA
TGTATACAGCCATAATGGTTTTACAGATACATATATGTACAGCCAAC
ATAAATATAAGTTATCAAATTACAGTAAGTTCTGACTTAATGTCACTAGG
TTCTTGAAACTTTGACTTTAAGCAAAATGATGTACAGTGAAACCAATTT
TACCATAGGCTAATTGATATAAAGATGAGTTAGGTTTTTGGTTTTTTTTT
TTTTTGACATGAAGTCTCGCTCTATCGCCAGGCAGGAGAAGAAGAGTTAG
GTTTTACAGCATGTTTCTGGTCACAAGAACATCATCAAACCTGTAAATAA
AGGCACAAAACACTTCTAATATTAATATCAAATAAATATGAGTTATAC
AGAATTTAAGAAAGATTAATAAAAACAAGTAAATCATTATTTATGGGAT
TTTTGGTAATCAGTGAGTTATGTGGTCATAGTGAAGTGGGTTAAGTCAA
GAAATAAATGTTTGCAAAACAAAAATTTTAAAGATCCTCTCCTACCACCA
CACAAAAACAAGAAAAACACGGTGGGCTCGCTAAGCACTTTTGTACCACT
CGTATCTTATGCGTTTTGTATGATTATTGTAAATGCTTTATGATAATTTTT
AGAGACAGGGTCTCACTCTGTGTCTCAGGCTGGAGTGAAGTGGTGCAATC
ATAGCTCACTGCAGTCTCAACCTCCCGGATTCAAGAGATCCTCCACCTC
AGCCTCCAGTGTAGCTAGGACTACAGTTGTGTGCCACCATGCCATCTAT
CTTCTTTTTTATTTTTTGTAGAGACAGGGGTGTGCTTTGTTGCCAGGC
TAGTCTTCAACTCCTGGGCTCAAGCAATCCTCCTGCCTCAGCCTCCCAAA
ATGCTGGGATTTCCGACATGAGCCAGCAGCACCTTGCCAGCATTTTATT

FIG. 4 (30 of 61)

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TCATAATAATTATAAGTCATTCCCTTCATTTCATCTTACAACCCACTTGTTTC
CAGTTTCAGGATCTCGGGTGACCAGAACCTATTAACGTTTCACGCACAAGTC
AGAAACCAGCCCTGGACAGGACACCATCCTACCGCAGGGAGAAGTTACAC
ACCCACACTCACTCAGACTGGGACCATGCAAAGAACCTAACGTGCACCTTT
GGAATGTGTGTTCCATACCCACTAGAACAGCTAAAATTTAAAAGACTGAC
CATACTTGAGTGTGTAACAGGATGTGACACAACCTAAATCTTTTAAGCGCT
TCGCGCTAAATGGCACAGCCGCTTTGGAAAACAGTTGGCAGTTTTTCAAG
TTAAATATACCCAACTCTATGATCCACTTCTCAACAATCAAACAAGAGA
AATAAAAGCAATGTCTACACAAGATGTATACACAATGTTTCATTGCAGC
CTTAATTATACTAGCCCCAAGTTGAAACAAGCCAAATGTCCATTACCAGA
TGACTGGAACATACAAATTGTGGTATATTGATACAATGAAATACTACTTA
GTAATAAAAAAGAAAGAGCTATTAACATAAGCAACAACATGGATGAATCT
GAAAACAATTATGCTAAGTGAAAACAGCCACACAAAAGTTACATACTGTA
TGATCACATCTACATAAAATTACAGAAAAGGCCAACTAATCTATAGACAG
AAAAGCAGATGAGTGGTTACCTAGGGATGGGGCAGAAGGGACGAAAGGAT
GGATTGCAAAATAGCACAAAATATTGGAGGGATGACAAATATATTTCATT
ATCTTGATTGTGGGGATAGTTTAAATGGGTATATATAGAGATCAAAGCTCA
TCTAATTATACACTTTTAAATATATGTATTTTCATTGTGCATCAGTTATTCA
TCAACAAGACTATAAAATAATATATGCCTACATACATTTTTTAAATATTCA
AAATCTCACAGTTATATACATAAATGCAACTGAATATGTATTTCAGATGTT
TTAACAAGCAGAAAGGACTGATTAACTCATGACAGCGGCTGTTTCTGGG
AAGGGTGTAAGAGACAAAGAGATGGAAAAGAGGATGAGAGCCAGAAGAGAC
CCTTGTAATGTTTCTTTCTTTTAGTAAAAATATATTGACAGTTAAAGCT
GAGAGGTGAGAATAATAGTCTCATGGCTTTTGTGTCTTAAATTTTACA
AACTAAGTGAAATGGGAGAAAGCAAAAAATAAACTTAAATAAATGTTAT
ATTGCCCAAAAAGAGATTTTAAATGGAGGTTAGACACATGAGACTTACGT
TCTCAAAAAGTAGAATCTGCAGGGAAAGTTTAAACAATAAAGAATTAA
AATCTAGCTTCTACCAGCCCAAAGCCTAAATGTTCTGCTTTATTCTTCC
TTATTATAATTCATAGGTAATATATTTTATGTTTGCAATGAATGCAGTG
ATATTAGATCTCTAAGAGGTGCTAAAAATGAAAAGTACATATTCCAATTT
TTCCCAATTTTCTTCTCTTCCATGAATGAAAAATATACATATTTGATG
ATTTCCAAGTTTATACAACCGATCTTTCTCTTAGTTTTCTCTTACCAAAT
TCCCTCCCTCACTCAGCCACCAGCCAGTCCAAGTGTGCTACCTGCACAGC
AGCCCTCATACCATCCACACTCTCATCAGGATCCTGCCTGACCTGCGAGG
AGCAGCAGCAAGAAGGAGACAGAACCTCCACGCTGAGCATCTCAGGGCTT
TCTCAGAGACTCCAGAGACCCTGATAGGGACAGAGCCTGGCCAGCAATC
CATGCTGCCAGCTGTATGATTGTGGGCATGTAAATCTCAACTGAAAATG
GGTGTAAATAATAACATGTTCTTCCCAGAAATGAGCTTTATGAAGATCATAT
AGCTGTTTTGAACTCAGACAAGCACTGGTAGGAATACAAACAGGGGAGCC
AACAGCCTATAAATAATACTTTAAGAAAGGGCATGAATGTAATTACTTAG
GAACAAAAGGCAAAGTGAGAGATGCCTAGGACTGAGCTGGACAAGCTGC
ACCTTTTAGTGGCTCAGCCCATGGGCTGACAAGGAAAATGGAGGAGCTAC
CAAAGAAGGTGGAAGGATTCTGGGAGAGTGGCCCTCACCTGCCCAGGGC
AGGGCTCAGTGGGAGAGAGGGAGATCTGTTATAAATGCTGCCAGGAGGTC
GAGTCATGTGAGAATGTCCATGTGAAAAACATCCACTGTGTGTATCTAAAG
AGAGTGGCTGTAAAAAGGTCAGGGTCAAAGGTCTTATTGTCTCAGATGT
TATCTGCATGCATTGTCTCAGCAACAAAGAACTAAGGAGCATGGACACA
AAGGGTTAGGTTGAAGCAAAAATTTAATAAGTGAAAGAAGAAGGCTCTCT
GCAGTGGAGAGGGGAGTCTGAGTGGGTTGCCACTTTGACAGCTGAATCCA
AAAGCTTTTATAAGAACTCTTCTCATATCTGCAGCTGTTTGAGTAACTT
CTCTTACCTATAAACTGTCTGTATAACTCTCCCTTATCTATGCAGCTGT
GGGATGTCTCCAGGTAAGCATAAAGTGAGCTTCTCTTGTGTTGTATAACT
GTGGGTTTGTGTTTAGGCAAGCCCCCATCCCTCCCTGTGTAAGCTCCCAT
GGAGCCCACCATGTGCATATCTGAGAAGTGAGGAAGCTTTCTCTGGGAG
CTCACTGATCGTACAAAGAACAAGAGGCTTCTGTGCCGCTTATCTATTCA
GGTGCAGCCTGAGTTTTCCCAGGCTGCTCTATTTTGCCTGTAGCTATG
ATTTTTCAGGCAGGCTGCTTCTCTGAAGACTAGCCTTAACTGTCTACCTA
TCAGATTTTTCTTTTCTTCTCCCTCAGCTGGTTCCCTCACCAAGGCTG
AGCAAGTGAAAAGGAGGGCACAGGGCAGGCCAGTAGTGAGCAGCAACAAG
GAACCTAAGACAGCAGAAACCACTCTTACACCTGGGTTGAAAGGGGTGGG

FIG. 4 (31 of 61)

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GAGCCAGGACTACAGC1 AGGTAAGAACATAGGTAAAGAGATACTGTTGT
TGTGTTGTTTTAACTATGAGAAGCATTGAGCTTTAAATTTCTACAGGAA
GGATCCAGTTTCAGACAGGAGCACCAATATTCAGAAGAGAAGACATGGT
GTAAAGGTCCTGGGAAGGCTGAGAGGATTGGGACTCAGAATCCAGAGCAG
AAGCCGCTCTGTGAACAGAAGAAGGACCTCCCCAGTGTAGCAAGAGGGAG
GGAGGAGGGACAGATGCCAAGATGGTTCAGGAAGAAGGTTTGGTGGTAAA
TGTGAGGCTGTGCTCACCTGCTGGCTTCAATTTCTCTTTAAATGTGAG
ATGGAATCATTGATGAAGGCCATGCCATGCAATGAAATGGCAGTCTGAG
GCATGGAGCAGCTCCAGCTTAGCCCGTGTTTAGGGTAATTATGGCTCAA
CCCAGGAGATGAATATGACTAGGGAAGTGAAGTCCAAAAACAAATGGTC
TCAAGTTGACTGTGAGTCTTCTGGGAGGCTGAGACGACAGGTGGGGTTGA
CAAGGGAAGGGGAACCCACCTGCTGAAAAACATCAGGCTGTTGGCTGGG
GAGGGGTGAGGCCTGTGTTGTAGAGATGGATGGATGCCTAAAGTTGGGT
AAGGTTTCAACTCTACCTCTGCTGGGTGTGAAATAAACAAAGACCACC
CAAATGAGAACAAACAAAGACTATTTATCCAGAGCTTGCTCTGACAAGG
AGTCGGCAACCATCACTTGCTTGGCAGAGACTCAGAAGTAAGCAGGGGAG
AAAGCCTCATAGCAGAAAGAAGGGAAGTCTTCATGTATGCCCTGAGTGGC
AGCTGTAGATGTGGGTGAGTTGCAGGTGGCTAACTAGAAATGGGGGACTC
CTGTGTGATTGATTAGGAGCATGTTTGGCTTTCTCTGGTTGGTCTACAT
TGGAAGAGGGGAACAAAAAATTTAGGGCAGTTGTCAGTTATTAATCAAGTG
TTGGCCATTTTGTACTGACTGTTACAGGAGTGAAGTGGCTCCCTGGATTGT
TTGCTAGAAATAGTGGTCTTCACTTCTGCAAGTCTGACTTTCTGGTAAT
AGGCTTCTGGGTGGCTATTGTGGATAATAAGTGGGTTTCTTGAGCTGA
TTTCTGCAGATTGTGGATCAGAGTTATTTTATATAAACAGTCTGACCATT
TTCCACTGGCATATTCATCTTCCAAGAGCTGGCCAAGCTGCTGTCTTAT
CTGTCTCCCCAGCCCCCTCACTCTGGCTGTGAAATACAAGCCACTAGG
TGAGGAATGGGGACAATTGAAGACTGAAAGCTTTTCTTTGCTGGGTTGCG
AGAGCTGAGGAAAGAAATGACAACATCCAAGTGTCTGCCCTGGGCCAGTT
TTAGGACTGTAGTGGTAATGCAAGGACTGTGTGAGTTTATATTTTCAATTT
GTCTCTCTAACTAAGGTGGAAAAAAGAGGAAATTTGCTGTCTGCA
GTCTCTGCAAAAGTCTAACACTGTGCTTCCCAACATTGCAGCCATTAGCC
ACAGGTGAGTATCAAGCACTTTAAATGAGACTGGTCCAACTGAGATGTG
CTCTGAGAAATAAACACACAGCAGATTTCAAAGACCTAGTACATGCCCTG
ATTTCAAGCTATATTACAAAGCTGTGGTAATCAAAACAGTATGGCATTGG
GAAAAAATAGACACATTGGTCAATGTGACAGAATAGAGAGCCCAGAAAT
AAACCCGTGCATGTATAGTCAACTAATCTTTGACAAGAGTACCAAGAAAT
CACAATGGGGAAAGTCTCTTCAATAAGTGGTGTGGGAAAACCTAGATATC
CACATGCAAAAGAAAGAAATTAGACCCTTGTTATACACAAAATCTAAAT
TAATTCAAAATAGAAAAAGACTTACATGTAAGATCTAAACCATAAACT
CCTAGAAGAAAACATAGGGAAAGAGCTCCTTGACACTGGCATTAGCAGTA
ATTTTTCAGATATAACATCAAAAGTACAGGCAATGAAAGCAAAACAAGT
GAGATATATCAAACTAAAAAGTTTCTGCACAGCATAAACAAATCAACAGA
GTAAAGACATGACGTATGGAATGAGAGAAAATATTGACATCTGACAAAGG
GTTAATATCAAAATATATAAGTAATTCACACAACCTCAGTAACAAAAGCC
AAATAACCTGACTTTTTTTTTTAAATGGGCAAGTACCTGAATAGGTATTC
CTCAAAAGAGACATACAAATGGCCAAGAGATGTATGAAAAGCTGCTTAA
CATACTAATCATCAGAGAAATACACAAATCAAAACAAGATATCATCTCA
CACCTGTTAGAATGGCTATTATTAAAAAATGAGATAAGTGTGGCCAGGT
GTGGAGGAAAGGAAACCCCTTGACATTATTCATAGGAATGTAAATTAGTA
CAGCCATTATGGAGAACAGTATGGAGATTCCCTAACAAAATTTAAAAATAG
AATTACCATATGACCCAGCAATTCACCTTCAAGGAATACATTCAAATACT
ATCAGTATCTCAATAAGATACTTGCACTCCTATGTTTGGTTCAGCGTTAT
TCACCATAGCCAAGATACAGAAACAAGTTAAATGTCCATCAACAGATAAA
TGGATAAAGAAAATCAGGTACATATATATACAAATGGAATATTATTAG
CAAAATCCTGACATCTGAGATAACCTGGATAAACCTGGAGGACATTATGC
TAAGTAAATCAAAGCCTGACACAGAAAGACAAATACACATAATCTCAC
TTACATATGAAATATGAAAATGTTAATTTTATGGAACAGAGTAGAATGG
TAGTTGCCAGAGCCTGAGAGTAGAGAAAATGAGATGCTTGTCAAATCAA
TCATCACATTGAATATATATAATCTATTTGTCAATTAAATATTTTAAGAA
TAAAAAATACCTGGCACCAAAAAAAGAAATGCAAAATGTCTCAACAATGTT

FIG. 4 (32 of 61)

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ATATGTATTGCATTTTG. AGTGATAATAATTTGAATATTAGGTTAAATAA
AATATATTTGAAAAATTAACCTTCACCTATTTCTTTCCATTTTGTTAACA
TAGGTACAAAAAATAAATTACCTATGTGGCTCATGTAGGTGGCTC
ACATTATACTTTGATGACACTATACAGGCTGGTGACCATATATCTCTTAG
ACTAGTCTAAGTGATTTAACAGTGGTTCCAGAAAGATCCAGGTTTAACAC
CAATGAAAGGGCCAGCTGGCTTAGCCCAGCTTGTGTGGGAAATGTTGGGG
AGTGGTTTAAGACAGGGAAAGCAAACTTTTGATGCTATTGACTTTTGTG
AAAAATCTTTTGTGGCTGAAAAAACCAAAACATTATT
>Contig49
GCTCGAGTGTGTCTCTAAAGCCTTTCCCCCATTTGGCTCCACTATACGCAC
TCTCCTGGTTTCTCCTCCCTCTAGCCGCTGTCTTTGGTCTCCTTTCTGATT
TTGCTGCGTCTCTGTCCCTGAATGATTGCTTCTCCACTACGGGGTGAT
TTTGCTCCCCAGGGGACATTTGGCAATATCTGGAGAGGTCTATGTTGTG
TTTGAGGGTGTGTCTACTGCCATCTAGTGGGGAGAGGCTAAAGATGCTGT
TAATGCCCAGGACAGTCCCCATAACACAGAAATTATTCAGCTCAAAATATC
CATGGTGCCAAGATCAAGAAACCTGCTCAAAATTAGCATGCTGCTGAAG
GCCCTTCTCTTCTTTAGCAATATCTGCCTCCTTAGGGATCTTTTCTAG
TCTCAGTGGTTTAAACATTTAAATCCCAATTAGGCAATAAATTGGGCCC
CAAACCTCGTTAGTATAAAATGTAGAACTGTGTTATTAGAAGGCTAATAA
AATGACCTGGTGAGCATCTGCAGCTAGCCTCTGAGCAATTCTGGGGACCA
CGTGCAAGATAAATCCATCTGTTCCCTCTCTGTAATGTGGCGCTACCTTG
TGGCCGATTTTTCTCGGGTTAAATATCTCTGGGGATGCAACTTGTCTGTG
GTTAATGGCTGTGTGAGGCCAGCGCTGGTGATAAAGGAATCAATCAAGA
CAATATTGAATTTAGAAAGGCAGATTTATTTAGAGAAAAGGAGAGATACG
TTGCAAGGGAGCAATGGGCAATACAGCAGAGGGAAGGCTGTCTGCAAGA
GGCAAGGGCTACGTATGACGTAGGGCTGCTTAGGCTGAATGCTTGCAGAC
AAGATGCTTGCCTGTCAGGTGGGCTGTGAGCTGAGTCTTGGGTGCTAGTG
AGCCATTGGCAGCTGACCTATTTCTTGGAACATTGCTCCCTGCAAGCA
TTTTAATGTTAAACCGCCAGGTGAGTTTGAATTTCTTTTTTCTTTTTT
TTTTTTTTTTTTGCTTTAGTAGGACCTGCCGTTGTGAGACTATCTGAGG
TAAATTAGACACCTCCTGGTTAAGTCACCGCTCCAGTGACTAGGCAGG
GAGCTCTTCTTGAAGAGGGTGTGGGCAGTGGGTACTTTGCATGTTGTCC
ACACCAGGCGAGCTGCTGCTTCAGGGCCTTTGCATTTGCTCTTTCTTTG
CCCAAAATGCACTTCTCTCACTGTTTACATGATTTTCTCCCTCTTTTCC
TTTTAGTCTTTGCTTAAATATCACCTTCTAGGGAGGCCTTCCACACCAC
CTCTTCAAGATTTGAGGGTATGCACCCCCACCCCTAGCCTTCTTATCCCT
CTCCACTGCTTTCTTCTCAAAGCACTTGTTACGTTCAAATAAAATAGATT
AGTTACTTTATAGTTCTAATTTTACTATTTTGTGTTACTTCAATAC
CCATGTAATCTCTGGAAGGAACGTTTCTTTTGTAGTGTATTTCTAGCAC
CTAGAACAGTACTTGGCACATGGCAGGTGTTCAAAGTATTTGTTGATTA
TTTTCTCAAAGGGCATGGAGTCTTAGAAGTTTGAGAACACAGTTCTAAGC
ACAGCTGTTTAGAGACTATGGATGATGCTAATGGCTGTATTCCCAGTAGG
TGGGGCAATTCTCAAATTGACCTGGAATCCTTGAGATCTGGGGACAGTCA
CCAAGCACTGGGCTCTGTGGGGAGAGATGTGCTGGTTTTTAGAGAGGAGA
ATAGCATCCTGGGGGACTTGGCCCCAGGGCTTTCTGTCCCAATCTCTTC
CCAAGTGTGCTCCAGAGGCAGGAGGCTTGTCTGTAGCTGGTCAGTCTGT
TAAGTGTTCCTCCCATCTACACAGATGCAAGAAGGCTGAGAAAAGCA
AGCTGTGAGGTGAGCAGGGGCCCTGACTCCTCCCCAGAAGGCACTCAGAA
CTTCCATAGGGCAACTGGAAGAAGGTTCTACTTCTCACCAGGAGCTGT
TGCTGGGGAAAAAACAGCCTCAGGCCCTACCCTGTGCTGAGAACCTGAA
TCCAGTATCAGGTTCTCCAACAACTTGGATCCAGCTGACCCTCACAAGG
GGTCAGATGCAACCTTGTAGCATATGGAATGGCAGCAAGGTCCTTGTG
TGGACTATGCTTGAATCTAAATTAAGACAAGGCCTCAGAGGGGCTAAGT
GACATCTGTCTCCAAAGTTTACAGCTAGTGTGACTAAATCTTGATT
CACCCTCTCAGGTTTTACCATAATCCCAAAAAGGTTGAAAACAAGAAAAG
TTATCTTTGGGCAATTACCTCTTTCTGTTTCTTACCTACTAATGT
TCTAGGCTCACCCTCTGGTCTGCAATCTCACTGAAGTACAGATCCCTCA
TGGCCTAAAGGGTTTTACACTGGGTTGACTAGGCTCTCCATTGCCTGT
CCTACTGTCTAAGGCACCTCCTGGGTAGGTTGCCAGCGTCATTCTGATG
CTGCTGACTTTCTTCCAGCTACTTTTGAACTTGGTATCCATGGCAGA

FIG. 4 (33 of 61)

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TCCAGCGTGAATTGGGCTACTTTGGGAGGCTGAAGCGGGTGGATTCTCTGAGCTCAGGAATTCAAGACGAGCTGGTCAACCGGTGAAACCCCTATCTCTACTAAAATACAAAAAATTAGCCGGGCATGGTGGCAGGCCCTATAATCCAGCTACTTGGGAGGCTGAGGCAGGAGAATCGCTTGAACCCAGGAGGCGGATGTTGTCTAGAGCTGAGATCGCGCCATTGCACTCAAGCCAGGGCAAGAATAACAAGAACTCTGTCTCAACAACAAGCGAACATACGAAACAAACGTAACATCCAACTAGCTAGCTATCGCGTGCCAGTCATGACCCATGGTCAATAGATGTCTACAGCTCAGGAAGCAGCTGCACAATGCCTGCATAGACAACTCTTATGAAAGCAGAATGTCTGTATGTCTCCATAACACATAACAGTGTATGCTTTTATTATGGTCATACTCTAGCTGTGATGTACCTACGCTCTAATATCCAACGATAGTTTTCTTTAAATCATCAACATAATAAATGTCATGCTGTCTGTCCCCCACATGTAGACATAACTTAGCTGGTACATGGATAAGAAACCTATATTAGATAACCTTAGGCCAGGTGTGGTGGGCTCATGCCTGTAATCCCAGCATTTTGGGAGGCCGAAGCGGGTGGATCACGAGGTCAGGAGATCGAGACCACTGTGGCTAACACAGTGAAACCCCGTCTCTACTAAAAATACAAAAAATTAACCGGGCATGGTGGCAGGCAGCTGGTCCCAGCTACTCAGGAAGCTGAGGCGGGAGAATGGCGTGAACCCAGGAGGCGGAGTTGCAGTAAGCCGATCACACCACTGCACTCCAGCCTGGGGGACAGAGCGCAAGATTTCTGTCTCCCAACCCAAAAANCNANNNNAAATTTGCACCCAAATCTGACTAATTCCAAGGCCAATTCGAATTTAGAATCGTTATATCTCCCTGGTGAACCTGAAGCTTTATCTTTAAGGAGACACACTCTTTATGTCTACCAATGCTTATTGECTTAAGTCCACTTTGTGCAGATCAGCTGCTTTCTTTAATTAGTTTTTGTGTGTATATCTCTTTCCATCTTTTCTTTTTCAGCCTCTCCATCTTACATTTTAGATATATTTCTTTTCTTTTCTTTTGTAGAGAGCTTCACTCTCTCTGCCAGGCTGGAGTAGTGCAATGGCGCGATCTTAGCTCACTGCAACCTCCACCTCTGGGTTCAAGCAATTTCTCTGCCTCAGCCTCCCAAGTAGCTGGGATTACAGGAGCCACCACCAAGCCCAGCTAATTTGTTGTATTTTTAGAAGAGATGAGGTTTCGCCATGTTGGCCAGGCTGGTCTCGAACTCCTGACCTCAAGTCATCCACCACCTCGGCTTTCCAAAGTGTTGGTATTACAGGCGCGAGCCACCATGCCAGCTGATTTAGCTGTATCTCAAAAACAGCATGGGTTGTGTTGCTTCTTTATTACGTTTATAATGTAAATCATTTACATCAAACTCTAATACACCATGGACTGTAAAAACACAGCCATATTTTATGTATGAATTAATAAAAAAACACCACCAATTAGTTCCTGAGACACACACCTTAACAATATCTCTGTGATGTGCATAAATCAATCACATCAGTTTCTCTGCACCTCAAAATTTCTTCTCAATTTCTCAGAGATATGGCAATTTCTCTGGTTTTACATTCCAGAAGCAAGAAAAAGTACACAGCTTCTTCAAGTCATGAGTAGCTTCTTTTATAGCTCTTGGTGTTGCAAAAAAGATTGGAATTGCTTCACTAATCTAAATTTTCATTCTGTCTGTCTGTCTGTATGACAGTCAGAGGCACTCTTTGAAGACATTCTAAACAGCAATTAAACTCAAAACATGTAATGACAATGACACACAAAACCTCAACTGATGACCAAAATGAAGAGTTCAGCCAAAGTTGACAAAGCTGGCTGACAGAGCTTGTAATACACACAGCTTGGCATATGCCTCGCCATTTAGAGATGTAAAAATAGGAATAAATGTTTTCCCTTAAATCAATGAAATAGAGCAATTTGGACTGAAAACTACGACAGTTATAGTGTGTTTCTATCTATTATCTCATTTCTGTTCTCTCCCTTGCTTTCTTTTAGTTTGAAATTTTTCTATCATTTTCTTTCTCTCTACTAGTTTGAAACTTATGCATTATTTTTCTATTTTTTAGCACTTACCTAAATTTACTCTGTAATCCATGGATCCTTAATTTATTTAAAAAACTAATGTTAATGAGTAGCTTTATTTTCTCTCCATCTAATTTAAGGCCACAGAACACCTTCACTTACCTCAATCCTCTCCAACTTACATGCTTTAATGTCAATATGTTAATACCGTATACTTTTAAAACTTTCTAAAAATAGCATTATTTTATAGCATGAGTGTTCAATTTACATTTTTGCATATATTTAGAATTTCTTGTCTCTCGTTTCTTCTTCTATTTATAGACTCCCCCTGGGATCAATTTCTTCTTACTTGAAGTACATAGTTTGAAGACTGCCTACTATTCAATACAGTAGCCACTAGCCATGGTAGCTATTGAAAGTTTAACTAAGTAAAAATTGAGTAATATTAAAAACTCAGTTCCTTCACTCACTAGCCACATTTCAAGTGCTCAGCAGCCACATGTGACTAATGACTACTGTACAGCAAACATATAGAACATTTCCATCATGGCAAAGAGCTCTATTGATAGTGTTCATCCAGAGTTTTCTGTTCCAGGACCAAACCTGAGGGTTGGGCTGCTATTTCTCATGAGCCCAACAGATGCGAGTAGCTGAGCTGGGGAGGAAGAGGTTTTTATTTCTGCAACCAGTTACAGGGAGAGGCCCTGGAAATCATCACCAGGTTTCACTCAAAAATTATGACGTTTTCCAGAGCTTATATACCTTCTAAGCTATATGTT

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TACGTGTAAGTGTGCAT1 JACCTGAAGACGTAAGTGATTAACCTCTTTT
ATCTGTAACCTAAGGTCTGAGTCCGGAAGATCTTCCCCTGGAGCCTCAGTA
AATTTACTTAATCTAAATGGGTCCAGGTGCTGGGGTAATTACCTTATCT
TGTCCTCTGCTAAATCATGGAGGTTTGGGGAATTCCTTTAGAGCACCAT
TAACCTGTTTGTGTAAGGCTGGGAATTTCTCCAAACCCCATTAACC
TGTTTAATCCCAAATTTGGTTCCGTTAAAAATTCCTCCTTAATTTGTCCA
ATTTTAAAGGCCCAAAAAGGCTGGGGCAAACCTCTGAATGGCCTTTGTT
ACATTCACACCTTTGTTTAAAAACACCGGTTTTTAATATTTAACTTAACC
ATTTAATCTCTACTGAAACACTTGTATATAAATCTGCATTAATGAGAAC
TGGCCTGCGCCATATCTCCTTCTCAGAATATCTTAGGGTTGTGATCCCC
GTGTGAAGAGAATATATCTCTGGAGATCTCAATCTCTCTACCCCAAAAA
AATCTCACTCGGAGAAAACCTCAGACTCTTATCTCCACAGCGCTATCTCT
TCCTCTCC

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GCTTGTCTAAGATGGTGTCTCCTTGTGCTGTGCCTGCTTTCATCCTGGGA
TCTCCCTTCACCATCAGGATTGCCTTCACCTCATTCCAGTCTTGGATCTT
TCTTCTTGTTCCTTGAGTATTTTTTTTTTTTTTTTGTGCTGATTCCCTTCA
GTGGCCTCTTGGGAAAAGATGTGTAGGGAGAAAAATTTCTTTAGAACT
TGCATATCTGACAATATATTTATCCTATCCTGACATTTGGTAGATAGTTC
AGCTGGGTACAGAATCTAATTAATTTCTTCTGATTATAAGACATT
GCTCCATTTTCTTCTGGCTTCCAATATTGCTGCTGAGAAGTCTGACACCA
TTCAAATGCCTGATTTTTTCCATGTGATTGTTGTTTTCTGTCTGGAGTGT
TGTAGGATTGCCTCTTTATCTACAGTGTCTGAAATTTTCATGACGTAGGT
CTTCTTTCATTCTATTATGGTAGACACTCAGTGGGCCATTTAATCGGGAAA
AACATGTGTTCTTCAAGTTCTACAACTTTATTACTTCTTTTCTTGTG
TCTTCTCTGGTCTGTTTTTCAAGCCCGAGTCTCTTAGATCTGTCTCTAA
TATTCCTATTGACTTTACTTCAATTTCTAAGTCTTATCCTTTTGCTTTA
CTTTCGAGAGACCTGCTTAACCTTATCTCCAACTCTTTTATTGAATTT
CATTTCTTTTACTATATATTTTACTTTGAATACACCTCTCTCTCTCTC
ACATTTTCCCCCATAGTATTTTGTCTTCAATTGACAGTCTACTATCTTA
TTACTCTGGAGATATTAATAAGTTTTTTAAATTTTATTATTTTTATT
TTCAAACAGTGTCTTACTCTGTCACTCAGGCTGGAGTGCAGTGGTGTGA
TCATGGATCACTGCAGCCTTGATCTCTGAGCTCAAGCTATCCTCCTGCTT
CAGCCTCCCAAGTAGCTGGAACCAAGGCATGTGTCAACATACCCAGCTA
ATTTTTTTGTTTTTGGGTGGAGTCTCACTCTGTAGCCCGGTCTGGAGTG
CAGTGGTGCAATCTGGGCTCACAGCAACCTCTGCCTCCTGGGTCTGGTT
CAAGCAATTCTCCTGCCTCAGCCTCCTGAGTAGCTGGGATTACAGAAACA
CACTACCATGCCCAGCTAATTTTTGTATTTTTGTAGAGACAGGTTTCACC
ATGTTGGGCAGCCTGGGTCTGAACCTCTGACTTGTGATCTGCCCACTTGG
GCTCCCCAAAGTGTGGGATTACAGGCGTGAGCCACTGCACCCGGCCACT
AATTTTTAAATTTGTTAATAAGACGAGGTCTTGCTATGTTGCCCAGTATG
GTCTTGAACCTCGTGGGCTTAAGTAATCTTCTGCCTCAGCCTCCCAAAGTG
TTGGGATTACAGGTGTGAGCCACTGAATCTGACATTTTTTAAAGTTTTT
TTCTCTTTACCAAGTCTTTTTTCCCTTTCTGCTTTTTTGGGTGTTTTA
TTTTGATCTCTATCTTGCTAGAACTTTCTGGAGACGTTTAGTAATACTA
GATTTTTGAGAGTGGGCAACTGGAAAGCTGATTGGAACTCTGAATACAT
GGGTGAGGCTTGTGGCTGTGAGTGTCAATGCTTGTGCTGGCAAGGC
CAATGGGTTTGGGACCCCTACTATTAGTATAGGCTGATTCCCTGGGAAA
GGCTCTTTTGATCTCCTGCCTGGAGGATAAAGGCTGGCTACCAGCCTTC
TGTGTGTAATGTGAGGGAGAAGGCTGGAGTATCAACATCATGCTGAAT
CCTTTCAATGATCATCTTGTTTTTTAGTAATCTCCTACCTTAACCTCTGT
CTTCTGCTAGTATGGGAAAGATGACCTGAAAATCTAACCATTATTTTTT
CCCCATTAATATCATTTTATGATTATTCAGAAGTTAAATAATTGTCATGC
TGTCCTCCAAAAGACTGAATCAACTAGCAACAAATAAGAATTTCTCAC
AGCTCTGCCAGCATTTTAAAGAATAGCTTTATTGAGCCCAGGAGGTCAA
GGCTGCAGTGAGCTGTGATTACACCACTCTACCCAGCCTGGGTGACAGA
GCAAACCCCTGTCTCAAAAAGAAATTTAAGGAACAGCTTTATTGTTGTA
AAATAGACATACAATAAACAGAGCACATTTAAATTGTGCAACTTATAC
TTTGATATAACCCCTGTGAAAACATCACCACAATCAAGATAGTGAATATAT
TTATCACCTCCTGATACAGTTTAGCTCTGTGTCCCCACCTAAGTCTCATG

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TTGAATTGTAATCCCCAATGCTGGGGGAGGGGCTTTGTGGGAGGTGAT1G
AATTGTGGGGGTGCACTTCCCCCTTGCTGTTCTTGAGATAGTGAATGAGC
TCTCATGAGCTCCCCCTTCACTCACTCTCTTTCTGCTGCCATGTGAGGAT
GTGCTTGCCCTCTTCTTGCCCTTCTGCCATGATGTGTTTCTGAGTCCTC
CCTAACCATGCTCCTGTACAGCTTGCAGAACTGTGAGTCAGTTAAATCT
CTTTTCTTCATAAAATTACCCAGTCTCAGGTGGCTCTTTATAGCAGTGTGA
AAAGGAACTAATATACCTCCTAAGTTACCTCAAGCTTCTTCTTAATTCCT
TCTCCTCCCTTCTTCAATTGCCAAGCAAACAACCACCTGTTTTCTGTAC
TATAGATTAGTTTACATTTTGTGGGTTTTTTTTTTTTTTTGTAGACAAGGTC
TCACTCTGTTGCCAGGATGGAGTGCAGTGGTGGCATCATAGCTCATTGC
AGCCTTGAACCTCTAGTTTCAAGTGGTCCCTCCCACTTCAGCCTCCTGAGT
ACCTGGGACTACAGGGGTACACCACCACAACTGGCTTAAAAAATTTTTTA
AATAAAAATGGGGTCTTGTTATGTTTCTCAGGCTGGTCTCGAACTCCTCG
CCTCAAGCAGCCCTCCCTCCTTGGCCTCCCAATTGTGGGATTACAGGC
ATGAGTCATGACTCCTGGCCTAGTTTACATTTTCTAGAGTTTGTATAAA
TGGAAACATACAGAATGTATTTTTTGCAGGAGTGGGGGAGTGTCTATT
TCTTTCTTTCTTTTTCTTTTTTTTTTTTTTTTTTTTGTAGACGGAGTCTCG
CTCTGTCTGTTGCCAGGCTGGAGTGCAGTGGTGGCATCTCGGCTCACCG
CAAGCTCCACCTCCCGGGTTCAAGCAATTCTCCTGCCTCAGCCTCCTGAG
TAGCTGGGACTACAGGCGCCGCCACCAACCTGGCTAATTTTTTTGT
TTTTTGGTAGAGACGGGGTTTTTACCATGTTAGCCAGGATGGTCTCGATCT
CCTGACCTCGTGATCTGCCCGCTTCCGGCTCCCTAAGTGGTGGATTACA
GGCGTGAGCCACCGTGCCCGGCCCAAGTGTCTTCTATTCTTAACAGCTT
TCATGCAATCTTTTTTATTTTACCATCTCTGTGATCCCACTCCCAAAGG
TACTAGATGTCGATTGGTCTTAGGATCAGCTACCATTTGCCAACTGCT
TTCCAGCCTTCCAAAAATTTTTCTTTTTTCTTAAAGATACTCCTGTG
TGAGGCTCAGAACTCTGAATTGCTACTGCAATATGAACTCGGTGATGT
GAATGCCAGGGAATTGCCTGATTGATCAAAGAAATGTATCCCTTCTCCC
TCACTCTTGCTGTCTTCTCATTTGTTTTCCCATCCTTGTGGATTCGTGA
ATTTAAATATCCCTTTAATGTTATAATATTTAATGGCGTTTGGCGAAAA
GTACAGAATTAGGTGCAAGAGTGCATAGCTGTTATTTTTTTTTTGGCCTC
TGAGACTGTTTATATATGCAAGTTATTTAACAGAAAGTCTGCAGTGACC
TGAGATGTCAGGGGGTCTGATAGAGTACGTTTGAAGGCAGTTACTGGAA
AAAAATAATGCCATTTCTGGTTTGTACTTCGGTAAGTTCAGATGACCCAA
TATATTGTTTACATGTGGCATTTCAGTAAAAAAGTAGCTTCCCTCCCTTT
CTTCTTCTTTTTCTCTTTCTGCTTCTATAAAGCATCTGCTTTGGGAAA
CTTCTTAGGAGGAGAGCTTGCCAGCCCGTGGGTAATGGAGAGGTCTTGCA
GAGATAAAGAGATGCTCCCACTCAATGCAGGATGGTGTGGAGGTAAATG
GGGATACGTCTGGCATCACTCAGGAATGGGCCTTCTGGCAGGGAAGAGA
AGGGAGGGGAAAGAGGAAGGGAGTCAAAGATGAATTGCTGAATACGGGGA
TTCCAGGGCTGGAGCCAGGAAGAGAACTTTGGGAGGTGTGAACCTGGAG
GGCATCAGCTGATGAGGAGCAGCCTGAAGTCCGGGAGGACCTGTTTTG
GTGGCCAGGAAGAAAGTGCCTTCCACACACAGGGAGGCCACAAGGCTGAT
GGGCTGGGGGTTGGAAGGACAGCCCTAGGACAGGCTTGGGAAGCAGGCTC
AGGTAGGGACTGCGAGGTTCTTGTTGAGTCTTTTTATTCTTGGTCTTAG
AAAAATAGAATCCAAGGCCTCTTGAGAGTGAAGGTGGGTTGGGAGGAGGG
CAGATGGGGCTTAGGCCCAGGACACCCGTAGAGCTACTGCCAGCTGTCT
CTCAGGGACTCTGCTGAGGTCACTCCAAGGATCATTTAGCCTTGCTAG
ACAGTACTGACAGAGGGAAACCGTAGTATCGCACCCACTTCTTCTCTTTC
AATGAAAGTTTAAAGGTCACCATTTCTCTGGCAAAGGAAGTTCCACAAA
TATTCCATTTCCGGTCTTAGAAACAGCAAGGTATCAAGCAATTGCAAACT
TCCTGTGCTGGGGAATTTCCCAAGGAAGTAGGGGAGAGTTCTGGTGGAGA
CAAAGTGAATTCGAGTGATTAGTCAGTAGCAGTAGCAGTAGCAGTAGCA
TAGCAGTAGCAGTAGCAGTAGCAGTAGCAGTAGCAGTAGCAGTAGCAGC
AGCAGAACCAAAATTTCCCGCACGTGTCTCAGGCTCTCATTTGCCAACT
CAGTCTCTAAGTATTTTTTATTTGGCAGGAAAAATAAATAGCTATGAGTGA
AATAATTCAATAGACCTGAGCCTCCATCAATTTTGTGTTTAAAGGCCTGA
CTCTCTTACCTTTCCCTGGGATGGAAGATGCAATGTTCTGATGTAC
TGTCAAAAAAGAAGAACAGTGGGTATATTGTATGCTTGAGTTCCAGCCA
TTTGTACAAATAGATAGAGATGACTGCCATGTGTGTAGACTTTCTATAGA

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CTGTGTGCTAAACCCGAUCTGCCACTTCCAAGGAGTAGATGAGGAATG.C
CATGGTTCTGGGGAGCCCTACCCCAATTTGGGGCAGACATTCCAAAGCTC
ATTTTCTGTGGAGGGGGTTGATGGTTAAAGGACGGCCTGGGAGTAACTCG
TCTGTACTAGGGCCAGGAGAGTTACATGCTGCTTCCCATGTTATTCATC
ATTCCCCCATGTGAATAGCTATGGCGTGAGGTCCAAGGTTAGGGCCTTTC
TACCATAAATGGGGGAATAAAATCCCCTACCAGCCTGAGAAGTTTCTGT
TATAAAGAGGCTTTTTTTTTTGGGGGGTGGGGGAGCAAGCCGACTAATGT
GTTATTTCCATACGGTTTGTGTTTTAAATGTAGATGTCATATGCAGGAGAG
GTGGTGTAGTGAGTCACAACGGGATTAGAAGGACCAGTCCGAAAAGCAGA
AGAGGGTCAAGTTCAGGGCACTGAGGACTACTGCATTCACTGGCGTGAAA
GGCAGATGGCTGAACAGGAGGGGGACATTACATTGCTTGTCTCCTTGAG
CCTCGATTTCTCATCTAAAAAGAGGGTCATTTATTCACAGAACATTTAT
TAAACTTGTGCCAGGCACCGTGCCAGGAGCTGGACTAAAAATTAAATCCA
CCCCTGTGAGCTGCTCTGAAGGCTAAAAATATGAAGTATGTAAAGTAACC
AAGTGCTGTACACATGCAGCTATTCAATGACTGTGTGGGCATTGCGGCAG
ATTTTAAATTTCTTTTTTATTTCTTTCTTTAGTGAGAGGTTGGTTG
TTATTATTGTGCTGCTGTAACGTCTATTTCACTTGCTTTTTTGTGTC
TCCAGCCCATTCCAGGGCTGTCTAAGACACTTCTTATCACCTAAATA
ACCGGGGAGGCAAAGCGCTTCTTAAGAGATGGATCCAGAAGAACATGC
TGGTTTTCTGTAGAAAAAGGGGCTGTGGGAAGTAGAGATAAGAAGGGAAT
TGGCCAAGATGAATGTACAGAGCCTTATTTTTTTTTTATAACACAGCAAG
ATTAGATACAAAACAGGACAATAGCATCATCTGTTTTTATACTGGAAAG
GACCTCACTTTACAGGTGGGAAGAAATAGAGTGGAGAAGTGAAGAGAATG
GTCACAGAGTCAATCAGCATGTCTGCGTCAAAGCTGGGATTCCTCAATTCA
GGGCTCTTACTACAGTGACGTATGGCTAATATTTTGGCATTGTTTCGGGG
AAAAGCTGAAGCCCTGATGGTGTACGTCACTCTTGAGATAGTCTGTAGTC
CAGCAGGGAGGAAAGCAAGGAAGGGAGGTGGAGGCAGCATTTTTGGGTGT
AACATTTCTGTTCTTGTGTTTTGTGGCCAAATCATAGTGTGATTGGGACAAGC
CACTGCCTTTCTCTGAGCCTCCACTTTCTTTTTCTTCTTAAGAGGGAGGG
AATAGTAGAGTAAAAAGTAGTCATTTTATCAAACACCTGCTATTTTGGAGC
CATATTGCAAGTGGGTTGGGGGTTGAACACTTGGCTTTATTACCCATAGG
ATTAAATCCAACCTCGATACTGTGGCATTCCCAAACCTCAGTCTAATCTT
CTTCTCCATCAGCCATGCCCCACGACACCCTGGTCATATCTGATGTTGCC
CCTTGCACTTGCCCCCTCCTTATCTTTGCTTTCTGACCTACCATATGGCT
ATTGGTTGAAATTTCTATTTTCCAGGGCCTTGCTTAAATATCATCTCATC
CATTAAACTTTCTTGAACCTCCCCTTGCCCTGTTCTCCCTAATGTCTC
AAGCCAGAATTTATTTCTTTTGTGGCCAAGGGACTGGGTTTGTGACCTC
TCTCACGAGACTTAATATTGAGACCAAACGTCTTTAGACCTCACCAGCCA
GAGAGATGAGCATCTATGGAATGCAGGCTTTTGCTGGACTGTGCTGATGC
AGGGCCTCTGCCTTCTCCAGGGCCTCTCTGCTGTTTTAGGAATTTCCC
TCATGGCACAGTCCATGAGCTCAGGGTCAAGTTCATACATGTTTTACTT
CTTCTACTCTGCAATGGTCTTCTTGAACCTGAGGGTCTTAAAGCTGCT
CTGCAGTTTGTGGGGTGAGTAGAAAGGGGCTTTCAAAGTTGTGCTGTTG
TTTCCCACCCCAATAGCATGAAACACAAAGATGCTTACAAATAGCTGCCT
TGCTTTCTAGTCCCACTTCTCTCTCTGAGGCTTTAAAACAAGTCCCCCT
AGGTTGAGCTGGACTGGAGTTGTATCCTATCTTCATTATCTGTCTACTCT
CTTTCTGCTCTCTAGAGAAGATATTATATATGTGTGTATGTATGTGAAA
TATATAATATCCATATATAGAACATATATTGTTATATTTACATATACATA
CATAACATATGCATGTATTATATATACATATGTAGTATCAAAGTTGGAA
TTAAACTGTATATTTTGTAAATTTGCTTTTATTTGCATCTATCACTGTAAA
ATGAATATTTATCCATACCGTAAGATATTCTTCAATGTATTTTTTTTTTT
TTTGAACAGGGTCTTGCTTTGTTGCCAGGCTGGAGTGCAATGACCCGA
TCTTGGGTCACTGCAGCCTTGACCTCCCCGGCTCAAGTGATCTTCCCACC
TTAGCCCTCTGAGTAGCTGGGACTAAAGGTGTGTGCTCCACACCCAGCT
TTTTAATTTTTTTGTATTTTTTTTTTAAAGACAGGGTTTTGCCACATTG
CCCAAGCTGGTCTTGAGCTCCTGGGTCCAAGCAATCCTCCCACTTTGGCC
TCCCAAAGTGCTAAGATTACAAGCATGAGCCACCACACCTGGCCTCAATG
TAATTTTAAATGGCTGTATAGTATTCATCATGTGGTTGTACCCAAAATT
ATTTAACCAGTCCCAGTTTATTTCAATTTTTTTTACTATTTGAATAA
TGTTTTAGTAAATACCCACAAAATATGTACAATGGCTGGGCTTAGTGGCT

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CACCCCTGTAATCCCAAACCTTTGGGAGTCTGAGGCAGGTGGGTCACTG
AGGTGAGGAGTTTCGAGACCATCTTGGTTAACATGGTGAAACCCCGTCTCT
ACCAAAAATACAAAAATTAGCCGGGTGTGGTGGCACACACCTGTAATCGC
AGCTACTTGGGAGGCTGAAGTAGGAAAATCACTTGAACCTAGGAGGCGGA
GGTTGCAGTGAGCCGAGATCACACTACTGTACTCCAGCATGGGCAACAGT
GAGACTCCATCTCAAAAAAAAAAAAAAAAAAAAAAGTACAATTTGTTG
TACCTCCCTGATTATTTCTTTTAAGTAGAATTTCTTATAATTTTTTTTA
TAAGTAAAATTTGAATCAAGGGAGAAGCACCTGGAGTCCTTCAGATACC
TATTGCCAACTGAACCTTTCTGTTCAGGTTTACTACATTGAGCCTGAC
TCAGGGTTTGGGGAGTAGAGGAGGGGTGGAGGCAGAGGGCCTCTCCCTG
TCCCCACAGACCTCCCTTGGTGAGGTCCAAGTCTGGACAGGTGGAGTGTG
GCATTGCACCGTCAGGTCTCTGCTTCTGTAATTCCTTAAATCCATCCAG
TGGAGCCTCATTGTTCAAGTCTTTTTTTTTTTTTTTTTTTTTTAACTCCC
CTGAAGACGGAGTCTCACTCTGTGCGCCAGGCTGGAGTGCAGTGGCACGA
TCTTGACTCATTTCACCTCTGCCTCCAGGTTCAAGTAATTCCTGCTGCC
TCAGCCTCCTGAGTAGCTGGCACTACAGGCGTGATCCATCACGCCCCGGCT
AATTTTTTTTTTGTATTTTTTAGTAGAGACGGGTTTCCATGTTGGCCAG
GCTGGTCTCGAACTCCTAACCTTGTGATCTACCCGCTCTGCCTCCCAA
GTGCTGGGCTTACAGGTGTGAGCCACCAGGCTGGCCTCAAGTCTATTTT
TTAACTCCAGGAGGCTGGTATTTCAGAGGGATTAGGGCTGGCAGAAGGGC
CTCAAAGCTTTCAGGCTGGGGAATAGGCTGCAGCCTGGTTCAGGGTAA
CCCAAGTGATTTTGGTTCCAAAGGACAGGAAAAAAGTGATTGATATGG
AAGTTGTCAAAGTGCAACTGTCAAGACATTAATAATTTAACTGATTGTAAGGAAAA
TAATATACAGTAGACTTGTGTTAAATATTTAACTGATTGTAAGGAAAA
AACCAGACGCAGTTTTCCCTACCATACTGTCAACACCTCAACACTGAG
TTCTTCTGTGACCTCTAGTCACCGAAATGCTTGGGGATTCTCCACCAC
TAGTCCTCCAGCAGCCGACACCAGTTGGGTGTCTAATTCCTCCAACAC
TATCTACCTGGAGTTAGCGTTAGATCCACAGGTTGAGGGCTCAGTCTCA
CAAGACTGCCTCCCACTTCAGGTGCCAGTTACAAGTGGTAGGTTGTCAAC
TATGCTTCTGACTGATGGCTATAAATCTGGGTTTGGTTCCTCGGGTTCC
GTGAATTTGCTAGAGCAGCTCACAGAACTCAGGAAAACTTAAGTTTAC
CAGTTTATTCTAAAAGATATTACAAAGGATACAGATGAACACCAGATGAA
GAGATGCGCAGAGCAAAGCATGTGAGAAGGGGTGTGGAGCTTCATGCCC
CTCTGGGGCACCACCCTCCAGGAACCTTCATGTGTCCAGCTATCTGGGAG
CCCTTCCAAACCTGTCTTTTTTGGGTTTTTAAGAGTGGCTTTATTACAT
ACACATGATTGACCGAACCTTGGCCATTGGTGACTGACACAACCTTCAG
CCCCTCCACTCCCTCCAGTGGTTGGGGAGTGGGGCTAACAGTCTCAAGTC
TCCAATCCTGCCTTGGTCTTTTCTGTGACAAACCCATCATGAAGCTACT
GCATTGGGGCTGCCAGCCAGCAGTCATCTATTAGCATGCAAAAGACACTC
TTATTATTCCAGAGATTCCAAGGGTTTTTAAAGCTGTATGTGAGGAAAC
AGGAGATGAAGAACAATATATATTTTCAACATCACACTCGTTGGGGGA
ATTGACAGGATAGCAAACTGATTAAAGGAGGATAGGAGAGACTGAGATA
TATATTTCCATATATATATATAGAGAGAGAGAGATATTTCCATATATA
TATATAGATCTAGAGAGAGAGAGATAGAGAGAGAAGAGTCTTTCC

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ACACATTTGGGGGAGCAGTTCCGGAGGTACAGCCCGACAGGAGATGTGA
GAAGATCGTGGTTANTGTTCCCTGGTCCAGAACCCTCCAAGTGGGCTT
AAGTAGGAAGGGTGGTGAGCGGCAGGTAAACACACGTCAAAGGCAGTCTT
CCTCTCTGAGGGAAAACACTTGTATAAGCATTGCAATCAATGGGCCTCTT
TAATTATGTGCCAGTGGCAAGAGCGGGTGTGAACCCAGGGGCCTGCCTC
AATCCGGGGCCTTTGAGGCAGAATAAAGTGGTCTCAGGTTGTTGGCATTT
CCTTGCCCTTCCACCCGAAGCAGACACAAATCCTCTCTGGAGGCAAGTTC
CCCAATTCAGCCAGTACAACCTCCACAGACTAAGATCAATCATGTACAAG
CTCACAGACAAAGGTCAACCAACACACAGAGCAATAAACAAATTCATGAG
TGACGTGAATGAGAATAAACAGAAACAATAACCACCAGCTGGGATGCTCT
AAGTCTTCAGCTGTTAGAATTCCTGAATATAGAATAAACTGCCACAATG
GCAACATGCATCTAGTACTTACTGTGTGCTGGGTTCTAAGAATTTTGCA
CATTGTGCCAGATACCGACTCAGCTTCACACTCACCCTCCTACTGTGCCC
TCTTAATTTGCACTAGATTAAAGGTAGAAAGGAAGAGGCAGCTATTCTG
TTCTTGGCTGTGCCTCTGGCAGCATGCAAAATGGGCAGTAACAGTGGC

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AGTCACAGGTAAGTAGC TTCTCACAGTGGGAGTTAAAGGCATGGGA
GAGACGAGCAAGGTTCTTAAAGGGACAGTGGCCAGTAAATGACCAGGGGC
TACTGGAGTGGCTGCATGGCTCTGTGGAAGCTCAGAGGAGCCTTGGGTCC
TGCAGGTGCAGTAGCAGCTTTCTGTAGTTCTGATCTCTGGGTCCACAA
TCTTCCCCGTTTTTGCTCCTCCACTTCTAATTTTGTAACTGACTTCCCTG
TGTGTACTTCTCTCTCTGATTGAAATAGCCAGACTGGTTTCTGTTTCCTG
ATAAGACATTGTCTGGTACGAACACAGTAACTCATTTAATCCGATATCTC
TATGAAGGAGGTACAATAATTATTCCTATTTTACAGATGAGGAAACACAG
CAGAAAAATAAAGTCAATTGTCTAAGGTTGCACATTTAGTCAAGGGAAGG
GTTGATATAACATATAATTATTTAGAAAAACATCTAAGGAAATAAAAGGCA
TAATTTAAAAATAAAACTAGGCAGGTTTAAAAAAATGAAGTAATCTATAA
GTAAAAAAGTATAATTGTTGAAATACATATCTTAGTGGATGGGTAAATA
GCTGAAGAAATGATTAATGAACTGGAAGGTAGTTCTGAGGAAATCAGAAT
TCAGCATAGATAGAAAAAATGGGAATTTACAAAAGTACACAGGAATTATA
AAAGAGGTTAAATTATAGGGAGGGTAGAATGAGAATTAACATTGGTCTAA
CTGGAATTTTGGGAAGAGAGAATAGAGAGAATGAACAAGGCAATATTTAA
AGAGGTGGCTGAGAATTTTTTCAAGACCAACACAACTATGACTTTACCAG
TAGAGAAAACAATGTACACTGAGGAGGATAAATAAATATACTATGAACAA
ATTGTAATAATAACTCAACAAAGACAAAGAGAAGATCTTAAAAATCAGC
AAAAAAGAAAGTCAGACTTAGAAAAGAAATGACAATGGCAGACTACTCAA
CAACAACAATGGAACCAAATTCAGTGAAACAGTATTTTCAAAATGCATA
TTAATCTATCTTTGAAGAATAAGGGTGAAAAGGGTGAAAATTGCTGCCT
TATACAAAATATCAACATTAACAAAAAGTAATGAAGGTAATATAAAATG
TTTTCAATAAACAACAACTGAGAGAGTTTACCACCAACAGCATTCTTA
AATGGACTTTTAAATGCAGTTTTTAGGAAGAAGGAAAACAATTCCTAAGG
AAGGTCTGAGATGCAAAAAGGAATTATGAACAAAGAAATTGTTAAAATTA
TAGGTGAATTAATAAACTGCCTGCATAAATGATAAATGACAATGATG
CTATTAATAATGAGTTGATAAGGATAAAGAAAAGGACAGAATTAATAAC
TAGAAAAACAAGCATGCTGGAAGGATTGAGGAATTACTTGAAGGTTAAAG
TTCTAGGGTCTTCTATCCTTCTAGAGGGGAGTCAATATATTAATTTTG
ACCGTCACTTACACAGTGAAAACTTTAAGGATAACCATAAAAAATAGA
AATAGAGAGTATAACTTCTGAAACAGTCAAGGGAAAAATATGGAATAAGA
AAACTGACCAAAAAACATCTCAGTCAATCAAAAAAAAAAAAAAAAAAGAAA
GAAAAGGTTCCGAAGGAGAAAAATCAAAGCATAGAAAAAGCGGGACAAATA
GAAGTGGAAGAAAGAAAAAGGTAGAAGAAACAGGTCCAGAAATATCACTGAT
GCACTAAATCACCATTAAAAGATGAAAACAAATGAACAACATCAAAAAAT
TCTAGTGACTGTAGTAGTGCTGATCAGAATAGGCTCTAAGATAAGATGCA
TTATTGTGAGTCAACTTGTGATGATGAAAGGTTAATTCACCAGAAAGAC
ACAAATTATAAATTGTAATCAAATAGTTTTATTTTACTTTATTTAT
TTATTTTTTTTGGAGACAGGATCTTGTTCTGTTGCTCAGGCTGGAGTGCAG
TGGCTTGATCTCAGCTCACTGCAGCCTCCACCTCTTGAGGCTCAAGCTTT
CTTCTGCCTTAGCCTCATGAGTAGCTGGGTCCACAGGCACACACCACCA
AGCCCTGCTAATTTTTGTATTTTTTGTAGAGATGGGGTTTCACCATGTTA
CCAGGCTGGTCTCAAACCTCCTGGGCTCAAGCGATCTGCCCCCTCGGCTT
CCCAAAGTGTGGGATTATAGGCGTGAGCCACGGTGCCTGGCCTCAAATA
ACTATTTAAGTGAAAACAAACTAGTATGGCACTAATGAAAAATGTATAAA
TCCATAATCGCAGAGGGATTTCAACTTACTTCTTTGATTATGTAAAGGT
CAACACAGACAAAAGACAATGACAAAACCTAATGCAATGAACACTTTTGAT
TAAATGAACATATATTGGATATGTACCCAAGAATTAGAGAATACATACTA
GTTTTGAGTTTATGCAGAACATTTACAAAAATTTAGTGAAGCCTAAATT
ATAAAAAAGTTGCTGTACGTAGAATAACACACAAACCCCTGAGTCCGGAA
TTCAAAGCCCTCCACACTCTCCTCTACCTTTGCATCTTTATCCTCCACCA
CACTGCAGTGCATACTCTGGGCTACTACTCACTGTTCTTGATTCAAATTC
CATGTTCTGTGAGCTCAAATCATTCTCTCTGCCTGGAATAACTACTTCAT
ACATATTCTGCTATTGAATTCCTTGCTTAGCACCCCATCTACTCCAAGAC
GATGTCCAGTTGGGGTTACTCCCTGTCCCATTCTTTGATTACACTTTT
TTTTTCTACTTCCATTATATTATTGATCACATCTGTGCCACAGTTTTTGA
CTTTGTGCTGCTTTTACTCTTTTCTAGACCCTGAGAGCTCCTGAAGGGT
TGGGTCAATTCCTTTTTTATTTGCTCATTCTCATGGCACAGTGAGTGCTT
AATAAATGGCTATTGACTGAAATTAACCTGTATCTAAATGGACATATTC

FIG. 4 (40 of 61)

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ACTTCTGGGCCATTTCATCTTTCTTTCTATTGGAACCAGGAGATGGGGAA
CCATAACAAAGGTAAGGTTGTGCCATGTGAAAGAACATGGAACCTTCCCC
TGAGGGCCAAAAAGAGCAGGGAAAGGTGCAAAGACAAAATCTTCCATTT
TTAAACAAATGTAAGAATGTGGTCCACCTCATGCTCAGGTGGGACTTTATC
ATGACGTTATTTTTGGGGACTTATAGCTGCATCATTTACCCCATATACAT
TTACCTTTAGTGTAGGGAACCTGAGGACAGGAATTTTGTGTATGCAGACTC
TTGCTAATGAGGCTAACACTTGGAGAATTTTTATCATGCATTCAAGAAGC
TTGTTTTACATTTCTTCATTAATACTTTAGTTGGTGGTTTAGCTTTAGTT
GTAGGCTTATCAGATATTTGGAGATATCTTCATAAACGATGGCTTTGGTT
TTAGAAGAGTTATTCTGAAGCTACTATTTCTGGCAATAATCAAACAGCAT
GGCCATTTGTTTTGTAAGGCCCTTTCCTAAATATGACGGTAAAATCTACG
TGTGGAAAAATGCTTATTCTTCTGTCCTCTATAAATGTGAATCTAGTTTG
TCTTCAAAATGAAATCAAGTGATTAAAAATGTAGTTTTCTAAGAAGATAAA
TGGAGCAAAGCACTCTGTGTTTCACAGTGTGGAAATCACTCATCCCTCA
TAAAACTGTCCCACTGATCCTGACTCACATGAATGAATAAAAATAAGAG
TTAATAACATCAATTTACATTTTTTAAAGACACTTTCCCATGTTTTAGACT
ATTGGTTGGAAAAGCTGGTAGGTGTACAATTTGTGGAGAGTTGGCTGTTT
TTGTCTGTGCTTTGTTTGACGTATTTCAAAGCCATATCTAATTTTGTGCA
GAATGGTCTGAAATCTACAAAAATGTTGAGTTGTGTAGTGTGGAGAAGTA
CGGAGCCATTTACTGAAAGGCTGGGGGGAAATGACGAGACCCTGAGATAA
GGCAGTAGTGGTGCGAACAGAGTGGAAGGGAGGTAGTTGAGATATGTTCA
GAGTAGAATCAGAATGGACATAGTGAACAACTGGATGCAGGTGGGGGCTG
AGGAAGCAAAGTTGAGGATAATTCTGAGACTTCTAGGTTGATCCACTGAA
GTTACATTATTCAACACCACAAGGAACTAGGGGAATGAGAAGGCATACT
GGTTTGCTTTGGAGTGGAAGGGCAGTGATGTAAGAGGAGTTAATGAGTTA
AAGTTTGGATATGCCTGAACTTCAATTTGATATGTGCATCTGATATACCC
TTGGGGTGACCCCTCCAGGCAATGGTTGAACATGTGTATTTCTTAGTAACT
GATAGGCATCACAGACTCACATCAGTAAGGAAGCAACAGCAAACCTTGATT
GGACGATATACCTGGAACCTCAGTACCCTATGACTGGAGCAAGTCTCTGTC
AGTGAAATGAGGATAAGAGAATCTTGACCTTGTGGAATATGTTGTTAGG
AATATATGTGATGAACAACTAGGATACTTCTACAGGGCTCCACATGTA
GTAAGGGCTTTATAAATGCTTGATAAATATTATTGTTGTAATTTATTTCC
AAAGTAAGATGCCACTGGAGGAATCTTTGGAACCCAAATTAATAACAAAT
AGGACTGGATGCAATGGCTCACACCTGTAATCCAGCACTTTGGAAGGCC
AAGGCAGGAGGATCTCTTGAGCCCAGAAATTCAAGACCAGCCTGGGTGAC
ACAGGGAGACCTTGATCTATGAAGAATTAATAAATTAACCAGATGTG
GTGGTGACGCCCTATAGTCCCTGCTGCTTGAGAGGCTGAGGTGGGAGGAT
TGCTTGAGCCCATGAGGTTGAGGCTGCAGTGAGCCATAATTGTGCCACCA
CACTCCAGACTGGGTGACAGAGTGAGACCCTATCTCAAATAAATAAATAA
ATAAATAAATAAATAAGTACAAACCAGCAAACACTAATCCTTTCTAGAGA
TTATTGAACTCTGGAGGGCAGATCTGAATGGAGCCAGCAGAGGGACCTAT
GGAGATCAGCCTGGCCCTGGACAGCACCAGGCAATGGGGTTGCTAGAGAG
GTAATGGGGTTGAACAGGGTTTAAGCCATGAGGTCTCAAGAATCCGTGAA
GACTCAGACTAATTTTTTTTTTTTTTTCATGAGGATTAGGTGTTCTTAGGA
ATTTCAATGAGAGCAGGGTTAATGAAGGAATGCAGGGTAGGAGAGCTGAG
GGAAGGCATCTGAGAGAGCCTGGCTTATGAATGGCTGCGTCAGTATGGCT
CACCTGCTTTCTTGTATCTACTTAGCAGATGATCCCACCCAGGCCTCC
AGGGCCAAGGTCAATTTCCACATAGTCATGGGCCCTTGAGGGCCTGGAGCA
GTGTAAGGAAGACAGAGTCTTAAGAAATTGCATTAACAGTCATGGTGCTT
GGCAAGTGTCGTCATCCTATGCCAAGCCTGATCTGAAGGGGTGCATGCTC
ATAGGTAGCTGCTGCCAAGATTACAGCAGCTTCTTCAATCCCAGATCCA
TGCTCTCCTATATTCAATTTTCCAGGGGTTCTGTCTTCGACAGTGATG
AGATGCAGAAATGACTTATTGAGTTATTCTCCTGATAGTTGCCAACTTTTC
CAAATGACAAATGGGGCATGGAGCTTGAGAGTGGAATGAGGCCCTAGGGA
TAGCGTGCTTAGGAAAACACTCCAGCCTGATGTAATTTGGGGGTACAA
TGGCATTTTCATCATCAAGACTGATGTAAAGGGTGACTAGCAGTGAGTTG
GGGGTGACTCGCACTGGGGCTAGGTTTCTGATTCTGCCTAATCCAGACAG
AGCAGAAGCACTAGTGGGCTGGTAGAGGGCCTCCAGGGCCTCACTTAATG
TCCTGGAAAAACAGCTCCAGATTGTTGGTTTACGTTCTGAGGACAAGCTT
GGGTACTACAGGATAGAGAGAGTGGTGGGAGATGCCGTGGCCTGCCCTGC

FIG. 4 (41 of 61)

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TGATGCCTGCCCTGCCATTCTCGGTGTGATGTCTCTGGGGCATCTTGCC
TTCCCTGCCCAGACCTGTAGTTCAGCTGAGGGCATGTGGAGGCCAAATGG
CTTCTTAGAGTGTTACTTTCTTGAACAGCTCTGCTGGGAGAACTGGAGG
AGCTAGCTAGTCACGGTAACCTGCAGCAGTCAAAGGATCGTCCCGGTGGAG
GTGGGGTGGAAAGGTAGAGAAAGAGAACATATAGCGTTTTCTTGGAGAT
GTGTGGGCATGTATAGAGGAAATACCCAATTCCTGAGCCTTGAGCCCTC
CAGGAAACCTTGGAATATTAGGTTAGTCATCCCCAAGGAAGTCTAAGAAT
TCTGGTCTCACCCATCTCCTTTAATTCACCAATGATCCTACATGATATT
AAGGAACACGGGCCAGTAACCTCCAAGCAATGGATGTGGTGGTGAAGTT
TGACCTCATGATGGAGCGGAGGTTGGTTTTGAAACCTAAGAATTTAATTTA
TTGTTTCAAACCTGTTCTCCACTCAGCGTTATTAAAGCATACATAATTGAC
ACATAAAATTTGTATATGTCTACGGTGTACAATGTGATGTTTCGATCTAT
GTATACATTGTGAAATGATTACAACAAGCTAAATAACATACCCATTCTATC
GTGTTTCAAAGGAATTAAGTCAAGCACAAAAGAGAGGTGCTGTTGAAGA
GTAGGGCTGCTCTATCTAAGTAGTATGTCTGGGGTGTCTGATCAGGG
TCCTTTTGTGCTAGTAATAAACAGCCCTTCTGGGGCTGCTCEACTTTC
CCACATTTTCTTCTGGAGCCTCCCTAAGAATTAGGACATGGCCACTTCT
CTGCATAGGCTTCTACTTCAACAAGGACAGGGCTTGTGCTGCCCCATGC
CACTTGAGTGTCCCTACAGCACAGAGCTGAGTGCACACTGGCTGAGTGAG
GAAATCCCCCAGATTAATCTTGGTTCTAAGCATCATGGCTGTATTTACA
CGTATATGAATTACAAATTACAGCATAGTCGAATAAGGATTTTGTGCTA
CAACTGGAATCCCAGATTATGCAAATTGGATAGTATAATATTGAAATTC
TAGGACTTTTTATTAGTTTTAAAAAATTATACAAGCTTAGAGTAAGAAAT
TAAACAGGTGCAAAAGAAATTCAGTGTGAAAAGTAAATGCTCTGTCTCTGC
TGAGAGACAGATATTGCAGCCCAGATACTACTGGGGTCAATAGTTTCCTT
TAAGCATGCCATTTTATGATGGTTTATGGGACTTACAGCTCAAGAAGCTTGA
CACTAGGGTTGATCTCAGAAAATCATTGTTGCAGGTATTAGATATGACCG
TCTCATAAAGATACACACACAGACACAGCGATTGGAGATATTCACTGGGG
CTTATGGGCTGCTTGTCTTTCTGCTCTGTGCCTAAGTTGGGCTCAGAGT
AGCCTGGCATCGGCTGTGGGGAGAATGCTGGCATGGGGTTAGCAGGAGCC
CACTTAACATGTCTTAAGCCACCTGGAAGAGTCTTCAAGGAGACCAGAC
TCCAGAGGCCCTAAGGAAGGAAGGACTTTTGCCGTTTTTAGGTATTCTA
GTCCCAGAGTTTAGGGAGGAATGGTTTGGCTTTGGGTGCTGTGCCCCCTT
ACCGAGTGGGATGGGATGTGCCCATGAGCTGTTGAGCTGGCTCTTGAGA
AGACAGCAAAAGCGGGAATAAGAGGTGAGGAAGCTGTGTGGTTGTAGGAA
ATCCCAGCAGAGGGCCTGGGGGTCAAAAGTGGTCATGGTAGTGACGGTGG
AGGCTGAGGTGGTAGAAAATCAGAGGACAAACCCCATGGGCTGCTGGTGA
TCTGACCCAGCTCCTATGCTCTCTGCTTCTTATTTAGGCTCTGTAGCAGC
AGATGATTGGCTGGTGTGAGAGCAGTGCACCTGCCATATCAGGCAATCCA
AGACAAGTCCAAGCTACGCTGGGAGGAAACCTGAAGGCAGCAGCAGGTAG
ACTGGCTGAAGACAGACAGGCAGGCAACTTGTCAATCAGATTTGTGTTTT
TAAGGACTTTTAACTGGGGAGCCCTCCATGACAGATCAGATGAGAGAGGA
ATCTGGGTCCGCCCATGTGTCAAGCTACCAGAGGGTCCCATCGGTGCTTG
GATCTTCTTTGAAGCTGGGTCTGAGGTTTGCAGGTAGAGGGTGAAGCTGGT
CAGAGGGACCTATTGCAGAGCTAACCACACCTTCCAGGAATGCAAGCA
CAAGCACCCACCGCGGGCAGGCGGGCAGGCACTTCTCCTTTTGCCACCA
GGACCTCACAGAGGCTGATCTGGCTCTGTGAGGTGGGAAAATGGGTTGTA
CTTAGTACATAGAGATAAAAGGCTTAGGAGGCCCTCCATCCTGTGACCC
TGTCCCCAGACCACAGGTGCCGGCAGGTGCTGCTATTTCAAGGCTGGGCC
TCAGTGCAAGCTTGTGGTTTTCTTGCCACCTGTGATGTCTCCCTAAT
GAAGGGGCTCTCCATCCTCTGTCTGCCTCTAGCAAGTGGAGGCTCTGGGC
CCTGGGCAAGACACAGGGGAAATGCCATCTGTTATCCAAATATATTTCA
ATGTGACAGGAAAGCTGTCTTTAGAGCACAGC

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GCATGTGCTCTACATTGATCCCAGGAGTTTGAACAACATTGCAAGACTG
GGCAACAAGCAAGACTCTGTCTCTACAAAAATAAAAAAATTAGTTGGG
CATGGTGGTACATGCCTGTGGTCCCAGCTACTCCTAAGTTGAAGAGGGAG
AATTGCTTGAGGCCAGGAGTTCAAGGCTGCAGTGAGCTATGATCACACCA
CTGCACTCTANCTGGGTGACAGAGCAAGACCCTGTCTCTAAAAATAATAA
TCGTAATACATTTTTTTAAAGTAAACAAAAAAGGTCACTTTCTCA

TTTTAGGGAGACATGAGACATCAGTGAATATATGTAAGATGTACACTGGT
TCCCTCCAGAAAGGCAGAACAACTTGAAGCAGGGAGGGAGCTTCCAGGTC
ACAGGTAGGTGAGAGACAAACAATTGCATTCTTCTGAGTGTCTGATTAGC
CTTTCCAAAGGAGGCAATCAGATATGCATTTATCAGTGTGAGCAGAGGGG
TGACTTTGAAATAGAATGGGAGGCAGGTTTGGCCTAAGCAGTTCCCAGCTT
GACTTTTCCCTTTAGCTTAGTGATTTGGAGGCCCAAGATTTATTTTCCCT
TCTACATCACTGTGGGCAGCTGACTAGGAAAGCTTTGTAGGACTGGTGGG
CAGTGTGAGAGCCAGTGGGGGGTGGTGGTCTGTGCCAATGGTAGCAAC
CACCTGTGAGGCTGAGTAAACTCATTTCCTCAACCTCCTCTAGCAGCCCCA
GTGGAGATACAGATGAAGCAGACTAGCGATACAACCCAGCCTGAAGTTTT
GTCTGGTGAGTGTAAATGGAATAAAAAATGGGAAGGGTGCTGAAGAGACCAG
CAAGAAAATGGTTGAAGAGATGGGGCAGAAAATTAAGCTGGATCAAAAA
GGACGGAAAAGCAGAAAGGGCCGATAGAGAGAGGGGATATCTATGGGTTC
GCGATTCTGAAAAGGACAAATCACTGGTGTCTTGAGAAGAGAGAGGGTGA
GAAAGCAGGAAGGCTGGAGGCTGTCTATCCAAGAGGCGGACATCTGTGAAC
ATGATTCCAAGAGTCACCAGACCATGGGGGTGGCCAAAGGGAGTGCCTCT
TCTCACCTCCTACTCTTAATTCCTTGTACTCAAGATAATAAGTTCCCAGA
AGAGAAGTACCCATATTTAATTCATCTGTGTCTTCTAGCAGTACTAAAA
ATATTATATGAAAGGTATCAAAACCTTTGAGAAATGTGTGCTGCTAAATTGT
TAAGGATGCTGGAAAACCTCAAGACGTCCCTGATCCTGAGCCTGAGTATGA
GCCTGTGGTGAGCCCAATGCAGGTCTCCATTGAGACAAAGGCCTCAGGGA
ACGGATGAGACCTAGGGACAGAGATGCATGCTGGAGCAGCATTCCCCATC
CCTACTGCAGCTCAGGCCAGCTGACTGCTTTATGAGTAAACGTTACCAGG
GAACACTTTGCAGTCTTAACACACATGCCACCTGTGACCACTGATCCCT
GTTGGGTGACCACTGACATCAGAGATTGATGGCAGCAATGAAGACAAGG
CTATCCTCATTAGGAAGGAAAGGAAGGAGGAGGAGGAGGGCAAACGAAT
CTTTCCTGCTTGTCAACCACGTCCATCTCTGTTAGGTGATTTCCCATGTG
TGACTTTGTTTATCTTTATAATAACTCTGAGAGGTAGGTCTTGATGTCCA
CATTTTGAACATGAGGACATCCAGCCAGGAAGTTGAGTTCTGGGGACATA
GCTGAGAGGGCAAAGCTACATATAAAACCCCTCTTTGTTTTTTCTGGCTTA
TCCACTGAGTGGCCCCCTGCAATCCACCAGCCCATTGTGTAAGTGCATACT
ATAGGTAAGTTGGCACAGGAGGAGTGGATGTGGGCGATTTGTACAGCT
CTCCAGGAACCTACACACTGGTGAGGAGGGCCAGGTATGTTCTTGACCAG
TCACAATCAAAGCAACCTCCTACTAATCAGGAGGCTTGGTACCTGGGGA
ATGCTATGTTGAAAGGTTCTTTTCTGGGTTTTAAATGATGGGTCTATTT
CCTTATTCTTAAGATTGCTTTTTTTCTGGCTAGAACTTAAAGAAATTTT
CAGTAAATTTCCCTTCCCTGGCACAAAGTGAGCTTGAATGAATTTCCCA
GGTGGCCTTGATACTTTAAATATTGCCTCCTATAAAATCAACCTTTAGA
AGAAGGAAGTCAAAGAACATGCTAGATTTCAAAAGGTTAATTCCTTGAA
ATCCAGTTATCTACAGGACAATGTTGTCAAAGAAAAAATTATTTGGCCAG
GCACGGCGGCTCATGCCTATAATCCCAGCACTTTGGGAGGCTGAGGCAGG
TGGATCACCTGAGGTGAGGAGTTGAGACCAGCCTGGCCAACATGGTGAA
ACCCCATCTCTACTAAAAATACAAAAAAATTAGCCAGGTGTGGTGGTGG
GCACCTGTAATCCAGCTACACGGGAGGCTGAGGCAGGAGAATCGCTTGA
ACCCGGGAGGAGGAAGTTGCAGTGAGCCAAGTTCAAGCCACTGCACCCCA
GCCTGGGCAACAGAGCAAGACTTTGTCTCAAAAAAAAAAAAAATTCAT
GATATTTTTAAATTCATGGTAAGGAAGATTTCAATCAGAACAGCACAGA
AGATATAGGAAACACTGCAATGGGACTTTGCGGTGGGGGAGAGAGATTGA
ACACAACATACATATACAGCACGGGCAAGGACATATTCATAGCCAGGAAGC
AGAGCAAAGATCAGTGGATGCGAAATTACTAAGAGGAAACATGAAAAATA
AGGGAGCTTCTGCCTAAACCCACCTAACCGGATCCTTGCTGAAGACAGGA
CAGGGTGATTGGACACCACTTTGGGGATGGTGGAGGATGGGGAATCCAGT
GAGATTTCAAGGGTGATCAGATATTGAACATAGAAGGTTCTTGCTAAAAA
AGGAGTTTCAAGAAAGTGTAACAAATGTGCCTGGGAGAAGGTTGAGGAGC
CTGACTAAAAATTTGGTCAAGCAGAGAATATTTGCCAAGATAATAGCTAAG
TCTTCTGACAAACAATAGATGCTAAGCCAGCAAGGGTGATGTGCTCAGAG
AAAGCACTGAGGGCTTATTTCTTTTCCCCCAATCTCCACTCAGTCAAGT
CTAGTCCCTTGTCAATGTAGCCATTTGTAAGAATGCAATCAGGCAGGGT
CCCATCTCCTAGTGACAGGACTGACTGAAGTTCTGCTGAAGAGAGTGGCC
TGGGGCTGACACCGAGATTTCAGAGTCTGGGTTTCGCCGAGAGCTCAGT

FIG. 4 (44 of 61)

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GTAGTGCCATGCCCTCTCTCCACCTGAACGCCAGTGTGGGCAGGAACAA
CTGCAGCTAGAAGTCTGGCACTTACGCTGGGGTCTAAGACCTGCCTGATC
TGCTAACTAGTCTTGTCCCTTGGCTATAAACTGACGTTGGCACCTGGCCA
GAAAGATGAGCAAGAGATCTCTGACACACCTTTAAGTCCCTGTGGAGTAG
GATTATGTTGGGAAGGTCAATTCTCTGACTGAGCAGCAATTTTCAAGAGG
AAGTCCCATGCCGAAGTGAGAGAAGGCAGGGAATCCTGCCTAGTCAGCTA
GAGCAAAACAGTCTGCAGGACGGGACCCAGGGATGTGATCCTCCCATCCA
AAGGCACTGAACTAAATGACTAAAATACTTTCCAGGGCTCACGTTCTTTG
AAGAATGGGGACTAAAATAAGACAGGAGCCAGCAAGTGAGGACTTGGAA
GGAGATGGCTCATCTGATCAGCCTCCACTCAACAATTTTAATCATCCACA
CTGGCATGGGGACACAATATGAATAAGTTGACAGGGACCTACTCTGATTA
AGCAGTGGGCTAGTGACAGAGACCTGTGAGTCAAGAGTGGACAGGAGATGA
TTTCAGACAGTGAGAACAATAAATTAACAGAGTCATGTGCTAAAGGGTGGCT
GGAATAACAGAGGAGTTTAAGACTCAAGAGGTCTGGCTGGGCGCGGTGGC
TCATGCCTGTAATCCCAGCACTTTGGGAGGCGGAGGCGGGCGGATCACA
GGTGAGGAGATCAAGACCATCCTGGCTAACGCAGTGAAACCGEATCTCTA
CTAAAAATACAAAAATATTAGCCAGGCGTGGTGGCGGGCACCTGTAGTCCC
AGCTACTCGGGAGGCTGAGGCAAGAGAAATGGCGTGAACCGGGAGGCAGA
GCTTGCACTGAGCCAAGATTGCGCCACTGCCCTCCAGCCTGGGCGACAGA
GCGAGACTCCGTCTCAAAAAAAAAAAGACTTGAGGGAGTTGTTTATT
TTTGTCTTTCTTTTAAGACAGGGTCTTTGTTGGGCGCGGTAGCTCACGCC
TGTAAGTCCCAGCACTTTGGAAGGCTGAGGTGGAAAGATCTCTTGAGCCCA
GGAGTTTGAGGCCACTCTGGGCAACATAGCAAGACACCGTCTCTACAAAA
AATGTGCAAGTTGAGGCTGCACTGAGCAGAAAAACACCGCTGCACTCTAG
CCTGGATGACAGAGCGAGACCCCTGTCTCGGAAAAAAAAAGAAAAAGACA
GGGTCTCGCTGTGTACACAGGCTGGAATGCAATGGTGCAATCATGGTTC
ACTACAGCCTGGAACCTCTGAGCTCAAGCAATTCTCCTACCTTGGCCTAC
CAAAGTTCTAGGACTACAGGTGTGAGCCACCACAGTGGCCTCAGGAGAG
ATCTTAATAATAAAGGACAAATTGCTTGCATCCCTTAGGGGCAGGATT
GACACATCCAAGGATCAGGCAGAAAGCCTGTGCGGAGTGGGATGAGCAAA
GAGAAAGGCTGAGAGTTGTGAAGAGGGAGATGCAGTGCCAGCTAGGACAG
GCCTTTTTGGGCTATGGGAGGTTTTTCAGAGGAGACCCACCTAACTAAC
CCATAACATTGCAGTGGGGACCTGTTGAAGTCATGGACTACTACCTGAAA
GCCAGAGAAATGGGAGGAGCCTTTCTCTGAGGAGGGACTCTAGTCCATA
GGTATCTTGCCACCAAATACATGGACAGGCCCTGGGGGAAGATGGTGGTA
GCCCAGCTGGAGGAAAACCATTTGCCACCTGAAGTACCCAGGGTAAGCC
ACCCAGGCACTGAGGCTGCACACCCATGCATGCACACAGAATCACACT
CCTTCCTATTATTCCTCAATTCAGGGTCTCAACACCCATTTTTTTGTT
TTTTGGGGTTTTTTTTTACATGTTTTACATTTTATTATTATTATTGTTGA
CAGGGTCCCCTCTGTTGCCAGGCTGGAGCAGTGCACTCGTGCAATC
ATATTAGATTGGTGCAAAAGTAATCACGGTTTTTGTCAATTAAGTTTTG
CCATTACTTTTTAATGATAAAACCACGATTACTTTTGACGCAACTTAAAA
GCTCACTGCAGCCTCAAAATTCCTGGTCTCAGGGAATCCTCCTGCCTCAG
CTTCCTGAATAGCTGGGACTACAGGCACATGCAATCCTACCTGGCTAATT
TTTTAAAAATTTTTTTTGTAAAGATAGAAAGTCATTTTGTGTCCAGGCT
GGTTTTCAAACTCTTGTCTTTGTGCTCCCTCTGCCCTGTGCAAGACCTTC
TGGATGCCCATAATGAAGACTTCAGGGAGAGGAAAAAGTAAACATAGGT
CCCTGATCAAGGGACCAGGGTTTATCGACCACAAACAGCATGCCAGATT
CCACTGGCAGTCTAGAGGTGCAATTTGCCCAAGTGTGTGTGGAAGGCC
TCTCCCTAGCAGTTGGTTTATACACCAGCCACAGCACAGCATATTCTCTT
AAATTGTGAACATTTGCAAAACTCCTTGAGGACAATATCATGTCTTGT
GTACTTTTGTGTTTTGTTTTCCCTTCCCTATGTACACGCGCGCATGCACT
CATGCACGACGCGCGCGCACACACACACACACCCCTCAAACTGAA
TGCCTGGTGTGCTGAATGGATGAATGGCTAATGTAAGTCATTCTAAAGC
TACTTTCTTTGGCATACCATCACCTTTGATTTTCATCTTTCTGGAACCTT
ATGTTCCCAGATGAATTTGGAAAGCCCTCAGGAAACATTTCAAAATTGCT
ATATGGGAGAAATGGGAGGGTCTCTCTAGAAATTTACCTGCCACAGGTAT
TTCTGGTAAGACACAGCAAAGGTGGCACCACCCATTCCTCGTTACAATGT
CAATGCCAGTCACCTTCCTGTCCCATAAACTTTATTAAAGGTGCAGAAAT
TCCCATGGAAGCAGGTGGACACCATCTGCTTCCAGCCAGCCAGGGGAGCA

FIG. 4 (45 of 61)

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AGGTGTCCACTGTGCCCTTGTGGCAGGAACTGCGCTTCTCTACTCTCCCA
CTTTGAGGCCTCTGGGGCTGGCCTGCTGCCTCCTCATTGACAAGGCTGCT
TACTGAGCAGTTTCATTCTGAGCTGGACATAGTGCTTCTGGTGAGTCTCTA
CTTCTATTTAACCCAAAGATATTCTTTCTAAGGAAACGCTTTCTGTGCG
GGGGAGGTTAGCTCCAGATGGAAGTCAAGAGTGATGGCATGGTAGCTCTC
ATCCGTTTGGGTGGATGATATTACGGAGCACCACCATGAGCCAGTCATG
GAGGTGAACAGTATATGCCAGCCCTGAATCAGGTGCATTGACAGCAAGGG
AGACAAGCAAACAAAGCTGAGGTTTGCTGAGGATGTTCAAGACTCACACA
GCACAGAGGAGCATCCACCACCCAGCTTGGGAAAGGACTTGTTATAGAGG
GGTGAAGCATGAGCTGAGTCTTGAAAGACTAGAAATTAGCCAAACTACA
AGGAGGAGAAGGAGTTTCCAGTCAGGAAGAAGAGGTTATGCAAAAGCACA
GAGACTAGAAAGAAATATCACATTCAAGGAAGTGCATAAGACAGGAAAGA
TTGATGCGTGGGATAGGAGAGGAGGGCAGGGGATTCCAGGTGGGCCCTGC
TTGCCACACTCAGGAGCTTGAACCTTATCCACAAAGGAGGTGTGGAACCAG
TAATGAATGGGTTTTGTGCAAGGGCTTCATGTCACCAGATTGCTTTTGTG
GAGATACTTCTGTGGCTGATATGTGAGGAAGGGATGGAGGAAGTTTCCGT
GGCAATCAGGAAAACCAATTAGCAGATGATTCAAATGGCCTAGGGGAAAA
GGGAGGAGGACTTGGACTACCATGCAGCAGCAGAAATGGAGAGAAATAAC
AGATCCCAGGCACTCAGGAAGCGCTCAGAAATGAGCCCTTCAAAGAACTTA
TGGTAGGTGATGGATGGATGGAGTGTGAGTCTTGGGATAGCATTGCCTGG
GAAAATACTTTCTAGTTGAGACAGGGAAGTGGGCCAGCAGAAATGGAGGG
CTTCTTCTTTTGTCTTAAATACTTTTATAATATTTGGAACCTTGAAAT
GAGCAGATATATTAGCAAAAAGCCTAAAAGGGATATTTTGAATCACTG
CTAGTTCTAACATATAACTTTAGCTTGCACACATCATCAATTAACCTTG
ATAGCGCTTCTGAAACTATCATCCCAATAGCAATCCTTGTA AAAACC
TATTTTGA AAAACGGGCCTTGTAGGATAGCCTCACAGATGTTTTGTGGTA
GATTTTCTAACATTCTAATGT CAGGGAGTGAAAGGAATCCCGTTAGAAGT
TGAAAAATTTCTGGAATCTCTATT CATGGTATTAAAGTTTTGCCGTACAC
AAAAGTTTAAACACCTTTACACAATCAGACTTCCTCATTTTACATTGCTCG
GTAATTAGAGGAAATCAGTCACCCAGAGCCTGGGTCTTAGACTTGACAAA
ATGCACCCAAACAAATCCTGAGTGGCCTTGCTGAGGACTTCTCCCAAGA
TAGAAAACCTCAGTTCCAGCCAACAAGGGGGAAGCAGCTGAAGAAGTGAA
TTAACAAAGTCTTGAAGGAAATGACCAATCATCTTTGATTGTGTAATA
ACCAGAGAGTAGAATACAGCTACGACAGACATTTTGGGAGAGAAGCATTT
TATCATAGCTTTTAGAAGAGAATATTTTTCAGCATCATAAGCACACAATT
CCAAGACAGATACTTTCAAGGGATTGTTTTGACG

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ATGTTNNGGTTTTTGGGACCCCATTCAAACTTCATGTTGAATTTTAATCTT
CAATGTTGAGCGAGGTCTGTGGGAGGGTGATGGATCATGGGGGTGGGT
TCTCCCTTGCTGTTCTCAATGATAGTGAGTGAGTTCTCAAGACCTGGT
TATTTGAAAGTGTGTAGCACCTCTCCCCTTCATTCTCTCACTCGTCACTG
CTCCGCCATAGTAAGATGTGTGTGTTTCCCCTTTGCCCTCCGCCATGATT
GTAAGTTTCTGGAAGCCTCCAGCTATGCTTCTGTACAGCCTGTAGAAC
TGTGAATCAGTTAGACCTCTTTTCTTCATAAATTACCCAGTCTCAGGTCA
TTCTTTATAGCAGTGTGAGAGTGGATGAATATAGTGCCATATGTTTGTAT
TCCAGCTACCCAGGAGGCTGAGGTAAGAGGATTGCTTGAGCCTGGGAGT
TTAAGGCTGCAGTGAGCCATGACTGTACCACTGCTCTCCAGCCTGGGTGA
CAGCGAGACCTTGTTTTCCAAAAA AAAAAACCCAAACTGTGTAAAATGTG
TTCATAAAAGTGTCTTGCTCCACACCTGTCCCTATATATCTTATTCTC
AGCCTCCGACAACCTACTTTATTCTTTCTTATGTATCTTCCAGAATCAAA
AAAAA AATCAAATACAAGCACAGTGAATGTATTGCCCTTCTCCCCT
CCCTTTTGTACATCAGAGTTAGCATATCATAAATACGGTCTGCATTTTC
TTCTTTTTCAGCTATCAGCATGTTTTGGAGAGGATTTTCAATTTCGTGCAG
ACAGCATGTATTAGTCAGTCCTTGCAATTGCTATAAGGAAATACCTGAGAC
TGCATAATTTATAAAGAAAAGAGGTTAATTGGCTCACAGCTTCGCAGGC
TGTTCCACAGGAAGCATGGCAGCATCTGCTTCTGGGGAGGCCTTAGGAAG
CTTTTACTCATGCAGAAGACAAAGCGGGAGTGGATGTCTTATATGGCAGG
AGCAGGACTGAGAGAGAGAGAGAGAGAGAGAAAGGATGCCACATACTTTT
AAACAACCATCTTGTGGGAACTCTGT CACGAGAACAGCACCAAGGGA
TAGTGCTAAACCATTATAAGAACTCCACCCCATGATCCAATCACCCTCA

FIG. 4 (46 of 61)

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CACCAGGCCCCACCTCCAACATCGGGGATTACAATTTGACATGAGATTTG
GGCTGGGACACAGAACCACAAATACCAGAGTGCTTTCTCATTCTTTCT
ATAGCTGCCTAGTATTCTATGTCCTTTACTTCATTTAGGCAGTCTCTTGT
TGATAGACACTTGGGTACTTCCAATTTTCCATTACAAATGATGTGCA
ATGAATAATTTTGATCATTTCATTTACATGGGTTATGTCCATCTGTG
GGATAAATCTCCAGGAGTGAAATTGCTGGATCAAAGGGGAAGTGCACCTG
TGATTTTCATAGTTAGCAAATTTTGTCTATAAGGGTCATATCAATTTAT
AGTCCACGCGTAATATTTAACAGTGGGGATTTCACGAGTTTGACCAA
CAAGGTCTGTGTAAACTTTTGATTTTGTCAATCTGATGGGAAAATAC
TAGTATCTCAAAGTGCTTTTAATTTGACTTTCTTATTACAATGTTAAGCA
TCATTTTACTCTGCCAAGATCAAATAGTATTTCTTTCTGTGAACAGA
CTGTTAAGATCCCTTGCTCTTGTGTTGCTGGATTTTGTCTTTTTTTT
CAAATGTTTTGAGGCAGTCTTTACATGTGAAACAAGTTATCTCTTATC
TGGGGTGTGAGTTACAACTACTTTCTCTGGCTTGTGCGCTTTGAC
TTTGCTTCTGGTGATTCCCGCAATCTGAAAGTGACTTTTGCATCATT
CATTCTTATACCCCATGCTCTTGTTCACGCTGGTCTCTAECTGAGGG
CTTTTTCTTTCTTTCTATCTGGGAACATTTTAGAGACAGGGTCTCA
CTCTGTCAACGCTGGAGTGCAATGGTGGGATCACAGCTCACTGCAGT
CTTGAACCTCTGGGCTCAAGCAATCTCCAGTGTGAGCTTCCAAGTAGC
TAGGACTACAGGTGCATGCCAGCATGCTGGCTGATTGTTTTATTATTT
ATTTATTTTTGTAGAGATGGGAGTCTCACTATGTTGCCAGGCTGGTCT
TGAACCTCTGGGCTCAAGCGATCTTCTGCCCCCTGCCACCCAAAGTGCTG
GGATTACAGGCGTAAGCCACCATGCCAGCCCCATGTGTGGAATCTTCTG
TTTATCCCTTTAGGCTTGATTCTTATGTCTGTTCTCTCCCTCTTCTG
CTACTCCTCTGTTCTTTATCTTACTCTACTTGTCTGTTACCTTGTTC
TGCTTATAACTAGCTGCCTCTCTATCTGAGGAGGAGTGTGACTGTTT
TCATCTCTGTACTCCAGGTCTTAGTACATAGCGCTTGCTCAACAGATGT
TTGGTGCATTGATAGATAAATCAATGGTAGCTGTTAATACCAGTCTTGAC
TCCCTGCAGTGCTTCAGCTGATCCTGTTCCAGATGTGCACTGAATATCTT
TCTGTTGAACAACAGAAATAAAGGGGATGGGTGAGGAGGATAGTCTTCGG
TGGCCAAGGATATTTGTAGGTACTTTGCAGCACTCAGCAATGAGGAGTGG
GCTTTAGTCCCCCAAGAACTCTCAAGCCCTGTTTGTCTTTACTGTTTCTC
TGTCAAATCCAAGACAAGTCAATGATCAGGAAAGACCTTTTTTTTCTTC
AGTGAAGTTTTATTTAGAACCATTGAACAGTATGATATTTGCTCATTAT
AAATATTTCCATTTAAATAATCTGAGCTTATATATTTTCAGTCTTAATTA
AAGGACTTGATTTAAAGAGAGCACACCAGTCCAAATTGAATTGATTCCAT
AGCTATTAATAAAGTAGGCTCTTTTACAGACACTGCTACTTCTTGCCCCCT
TTGAATAAATTAGACCAATGAATAAAACAACAACAATAAATAAATAA
ATAGGGAAGCGGTTGCTCATCAGAATGTGGGAGCGAATGACAGAGGGTTT
CTTAGAACCAATGTGGCCGTGTTTCTGTGAGGCGGCTTTAAGTGAGT
AGGAGAGGTGAGAGAGGCTGGCTCAACAAAAGGGCTGGGGATTGGCCCT
GAAAGGAGAGAGCTGACTGTCTGGCTGATGGACAGGAGATCCTCTTAGC
ACTACCCTAAGGCAGGCTGTTGGGCATTGGTGTAGACAACAGGAAAGTCC
AGGCTATAGCCGTACTCAAAAACCTTTCTGTTCCCTTTCTGCCAGCCCTA
GGGATTGAGTCCACATTCAGCACAGGACTCTCTGGGTACAGCTCTCTTA
GGAAGACACAAATTGCATGGTGAAGTCAGTTATATCTTGCCCGCTTTGG
TCCCTCCAGGAAGACGGGCATGTTTTCTGCTTGAGAGGTGCTGATGTAC
CAGTTGGGGAACCTGGGCAGACTCAAATTCAGCTTGTATTGATTCTAT
CTTGTTGAAGACAAATCGCTTTTCCATCTTCTTTGGGTAATTTTTGG
GATCTACACTCTGCAGCGAAAGAGAAAGAATTTTTGTGGGGCAAGGG
ACAAAAATGCTATGGGAAAGATGTTCTTTGGGTGGCCAGAAAGGAACT
GACGAGCAGGTACATGATCAGGAGCCCACTCCTGAGTTGTAAGTGGGC
CCCCAATTTCTGTGTGATTATTAAGAGAGCCCTTCTTCTTTCTAAAC
TTAGTGCCAAATGCTGAGGAGCATAATGTAGGTGAGAATTTTTTTTTTT
GGGGGGGTGAAATTAAGCTAGAGCTTCTTGAAGTACCTAGTTTCCAGGG
GCTTTTTATTGTATTTTCTTATGGTCTAGAATGACATCAACTGGAA
ATGAAGCTTTTGTGAGAAAGCTGGAGGTGATAGTGGTGGTGAATTTGGG
AGTGGAGTGGACGTGATAATGGGACCTTTAAGTCATCTATTTCCCAAGG
TGTCTATCAAATGAGAGCAGCCCTAACAATATATAATCTGTTGGGGTTGT
AACTATGGTAGGACATAATAACATCGGCAAAATGATTTAATTTTCTGCAG

FIG. 4 (47 of 61)

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CAGGATTGAAGGTTGCAAGCAGTTAAAAAATTATGTTAAATTTATTACAT
TAATGCAAAATTGTCAAATAGACCTGTTCCAGCTTTTCTAGGGATGGG
GGCGGGGAGAAGGTGGTTGTCTGGGAATAAGTGGTAGCAGGAGGCTGAGA
AGGGCTTCATTCCATAGCATTCACTTACCTCCAGCTGTAGAGTGGGCTTA
TCATCTTTCAACACGCAGGACAGGTACAGATTCTTTCTTGGAGGCCAA
GGCCACAGGTATTTGTCACTTACTTTCTTCTCCTTGACAAAGGACATGG
AGAACACCACTGAAGAAAGAGGGGGTCTTGTGGTTAGGGACACAGCAGT
GCAGGGTCACCCCAACCCCTAGGCCCATGAGTAGGATACATGTAATTTG
GTAGCCTCTGTGGGAACCCACAGTGAGGTTCTTGGCCTAAGACACAGGA
TAACTTGACTTCTCACAGACAATAGCAGGGTCATTTTGTGATTAGGGT
TTCCCTCAAAGGCCTGAGGGTTTCTCAGAGCCTCATAGCAGTAGGAACG
GAGAATGAAAGAGGGTCTACATTTTAAATGCTGAAGGAAGGAAGGAAGGA
AGCCATTGTGTCACTGGCTGGCAATGTGCCATCCACAGGAGCGGAACAA
CTTGATCAATGTGGAAGGAAGGAAGGGTGGAGGCTGTACTTCTGCCAG
AAATCAGGCACCGAAGTGTTCAGGAACAGAGAGTAGCCCATGGGAAGA
AATGGGAGAGGAGAGGCTGAGCTGGGAAAGTGGCTCCAAAGAGAGACAC
TCATTTTGATCTTCTCAGTCACAGCAGTGTCAATTGGAAGGCCCTGGGA
TCACTCTTACTACCCGATTCCAAAGAAACAGGATTTTCTTGGCCTGGCTG
AGAGCAAATAGCTTCCCCCTTGAGTGAGGCTGTCTTCAAAGTCAGCAGC
CTTAGTTGGCCACACTCCTGTGCAGAGGCTTGGCTACTGTGGCAGCATG
CCAGGCAGATACCCACAGCTAATGATGGGTTCAACCGACTTGAAACTTTT
GCCCCCTACAGCGGAGAGATATAAGTTCTGTGGGCGGTAAAATTTCCC
TACAAGGAACCACTGGCATTTGGGTGGGACGGATGTTGGGGCAAGGGGGG
AAGACTGGGGAGGGGGATGGACACATTATCGCTCCAGCACTCTTGTTC
GCCTCAACAACAGGAAGAGAGAACCCACAGGCAGTTAGGCCATGTCCATC
AAATGACCCCATATTGTGGAAGAAATTGACATTGCACTATGCCCAAGAGAC
TTGGGTGGACATGGTCTGGGAGTGCTTGAGCCGTCTAATTTCTCAGGGT
CACACTCCTGTTAACAAATGCACTGGCCAGTGCAATCAAATGTGCCATTT
CTAGGACCAAAGTTTGTATATTCTTTTAAATATTTTTTTCACTTGTGT
TGATCATTTGCCCTTAAATTAACCTTTCTACTTTGTTTAAACATGGAGAAT
TAGCAAGCTGCCAGGAAGCCAGGCAGGGAACAGGATGTTTCCATTTAC
CTTGTGTCTCCATATCCTGTCCCTGGAGGTGGAGAGCTTTCAGTTCATAT
GGACCAGACATACCAAGCTTTTTTGTGTGAGTCCCGGAGCGTGCAGTT
CAGTGATCGTACAGGTGCATCGTGACATAAGCCTCGTTATCCCATGTGT
CGAAGAAGATAGGTTCTGAAATGTGGAGCATGTTGTTTAGGTATAAAA
TCAGAAGGGCAGGCCTCGTGAGGCAAGGTGGCAAAATTTGATTTCTTGA
GGACACCTGAGCATATACGGTCAAAGTCTGATGACAACACCAGTAGGGAT
GAAGCTGGAGTGGGGTGGCTAAGAACACTGGACCTGACACTATTAGACA
TGGGTTCCAGCTTCAGGTCTATTACTGCTCACTGTGGCCGAGCAACAGAG
CTACTTAGGTAAATGGTGATGGTCATAACACTAGCCACAGGGAGGTTA
CGAACCTCTGGTGACAATGTAAGTGAAGGGCCCCCTGAGAAAGAGTGAGGG
AGTTGCAAAATGTCAGTAGCCATCAAGATCTTCTTAAAGATAGTTTCCAC
TAAAGAGATGATTGCTTTGGTTTCCAGCCTTCTTTGTTTTGTCTCCCGC
TGGGCCTTCTACCTTTAAAGGGCTTTGGCTCTGGGGGAATTGAGTTGGCT
TTGGCTAACTACCTTCTTCAAAGATGAAGGGAAGAGAGGTGCTCAGGTCA
TTCTCCTGGAAGGTCTGTGGGCAGGGAACCAGCATCTTCTCAGCTTGTC
CATGGCCACAACAACCTGACGCGGCCTGCCTGAAGCCCTTGCTGTAGTGGT
GGTCGGAGATTCTAGCTGGATGCCGCCATCCAGAGGGCAGAGGTCCAGG
TCCTGGAAGGAGCACTGCGGAGAGAGCGAGGGAGGAGCCTGGTGAGGTG
GTCCTGCCAGGAACCATGCTTTGACATCAGAGAGTAGAAAGCTCAGAGAG
GAGGAAAGGGCTTGAAAGAATCCCGAGCTTCTAAAGATCATCCCTCTCTG
GGCCAGGCGTGGTGGCTCATGCCTGTAATCCAGCACTTTGGGAAGCCGA
GGTGGATGAATCATTTAGGTGAGGACTTCAAAACCAGCCTGGCCAACATG
GCGAAACCCCTTCTACTATAAAATACAAAATTAGCTGGGTGTGGTGGG
GTGCACCTGTAATCCTAGCTATTCAGGAGACTGAGGAAGGAGAATCGCTT
GAACTCAGGAGGTGGAGGATGCAGTAAGCCAAGATTGTACCACTGCACTC
CAGCCTGGGCAACAGAGTGAGACTCTGTCTCATAAAACAAAACAAAACAA
AACAAAACAAAATAAAATAAAATAAAAGATTATCCCTCTCTGAA
GCTCAAGGAGGTTAAGGGTGTACTCAAGGGCACACAGCAGGTTAGAGGCA

FIG. 4 (48 of 61)

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GA CTCAAGACTAGAATG TGGGCTTTCTGACACCTTACAGGCTATTCTTTT
AGAATAAATCCCATTCTACTTTGTTTATCTTTTGTACATGCCCCACC
TACACCATACATGTATACCTTCTCTATATCTTTTGTATCCCTAATGCTG
TCACACTATGATTTGCTTTTTCATGCAGATGACCATAACATTTCCATTCTC
ACCTATGCTCACTCAGCAAGTATTCAATTTTCTACACTGTTCTTTTTTT
TCCTTTTTTCATAACACTGTCTCATAGGCATTCTGCAAATCCTGTGAGAGT
ACTTTTTGTGAAATGTTACCCTTTCTCTTATTTCAGAGAAGCTCCGTAT
TAAGGCTTCACTGAGGTTGCCTTAAGGCATGATAATGGTTCAAAGGCTTG
AAAGACAGTTAAAGAGACCTGTAAGTGCACAAAAGAAAGTTGAGCAGGAG
AGAATTTCTTGCCTGGAGCAGAGCCAAGCTACTGGAAGAGGCAATGGGGG
CAAAGGCCAGGCAGACAAGCCAATGGGCTCCTCCACAGCTGCAGCCAAC
AAGTTATGCCAGTCTTAAACCTTCTAAAGAAATATGTTTTAAACAAGATT
GAGGACTGGATTATGAGGCTAGGGGAGGCTATCACAACTGGAATAAAAT
AAAGCCAGAGAAAAGTGGCTGCCTTCCAACCTGCACAACCTGACCTAGCTA
GGCTGATGGCTGGGCCACCTAGGAAGGCTACTGAGCATCATATAAAACAG
AAGGGACAGCAGGAATATAACATGGCTCTTTGTAAGGATGAGTCTGAAAA
ATGACCATTTGCTGCCCAAATGCCCTTAGCTACAACCTGAAAATATTTTCA
AACTGGAGGTTGCAGGATGCTGGAATCTCAGAGATCATCCAGCTCAGCCC
TTTATTTTTTCAGATGAGGTCCAAAGCGGGTAAATGACTTGTCAAGGTCA
AACAGCAAGTGAATGGTTTTCTTTCAAGTCTCAATTCATCTTTTTGTTTA
TATCATCTATGCTTGTGTTTATAAGCTTCAACCCAGGTAGCAAAAACT
ATTCTACTCAAAGGGGTAGACATATGTTAGTTCTCAAGATCATCTCTTG
GTTTCAGAGTTTAACTCAAGTGATTGGCATAGGCTGAATCCATCTCTTAA
AAGGATAATCAAATTTATGTTGAAGACTTGGTTGTCTTCTACTATGAAA
TGGGAAACATTATCACTACTCCTCCCTGTCAACCAAGTGTGGCCACC
ACCACCAACGTTAGTGAGTGACTGTGGTGATATGATGACCAAGTGGCCAG
GTCAGCAAGTGGTGCAGCCTGTGTCTCACTGGAAGAGGTTAAAGTCTTTC
TAAAACAAATACCATGGCATCAAAGTGGCCCAAGTCCCTTCTTTGAG
CTTTCCTGTGTTAGAGCCCTTCTTGGGTTGGGAGTTAAACCCATAGTC
TTACCTTCATCTGTTTAGGGCCATCAGCTTCAAAGAACAGTCATCTCTCA
TTGCCACTGTAATAAAAAACAGGGACATGTCTCAATTATGTCTTCTAAACA
GGTTTATTTTTCTTCCCTGTGTACAAGACTTGACTGTTTATAAGAACT
GCAAACAGCCTGCCTCTCAAAGCTGCCTGAAACACCTGGCAAGTTTCA
GTGATATGCCGAGAACAGTCCAGAAGGCAGATTCTAGGCCTGGCAGGTGG
GCACCTTGGGTGCTCCCTGTGATCTTGAGGCCTAACCTTAGCCACAGC
AGAGTCAGCTAAATCTGAGCTCTCCCTCTCCCTCCAAGCCACACTTTC
AAAGGGATTCTTGTATTGTGGCTTGGAAATCTTTCTCCCCATTGCT
CTGCAGGAAGCCCTTGCAACAACACATCTGGATAGCCTCCAGGTCCCAAG
GCTGGAGGGACTTGTAAATGGGAAAGTAGCTTTAAATCAGATTTACTTGG
CACCCTGTTTGCCACTGAAAGAGGCAATTTAGGGGAAAAATCTGGTCTCC
AAGCACAGATAACACTCTACTCTTGAAAGAGGAGACCTGCTCATGTTACT
GGTCTCAGCGTCTCCACTGACCTGTAATAAGCCATCATTTCACTGGCGAG
CTCAGGTACTTCTGCCATGGCTGCTTCAGACACCTGTGTAAAAAGGAGAA
AATGAGTGACTTCCCCATGACGGCTACGTTTATGTGTGATTTCTCTCAGC
ATCCAGTGATGGCAGTCATGCAAAGAAATGATCTCTGAGTAAATGAATG
AATGTGTGAAAGAGAAGTCTTTGGGTCTAGAGAAAAGCATTTGCTAAAC
CAAACCCCACTAGCAATGTATTGGCTAGGAGAGCTGGAGCAGAGGCTTT
GACACTAACCTTTAGGGTGTGAGCTGTAGATAAGCAGTATCCATTCCCA
GAATATTTCCCGAGTCATAAGCATTATATTACCTGGCATTTTTGCAAA
AAGCTGAGAGAGGGAGGCAGAGAGGGAAGGAGAGGAGAGACAGAGAAAG
AAAGAGAGAGAGAGAGAGAATATGCATACACACAAAGAGGCAGAGAGACA
GAGAGACTCCCTTAGCACCTAGTTGTAAGGAAGATTAAAGTCATACTTGA
GCAATGAAGATTGGCTGAAGAGAATCCAGAGCAGCCTGTTGTGCCTTGT
GCCTCGAAGAGGTTTGGTATCTGCCAGTTTCTCCCTCGCTGTTTTATAG
CTTTCAAAGCAGAAGTAGGAGGCTGAGAAATTTCTCTGTTGAATACCTG
ATTTCACAATCAAGTTAAAGGAAAGGGGAAAGAGTATTGGTGGAAGCTT
CTTAGGGGAGGGGACTAATAAACTGAGATAATTTCTCTGGTTTATGGAAGG
GCAAGGAGTAGCAAACCTATGACACATTTGCAAATGTATCACCATGCAAA
TATGCATTTGTTTCTGACAATCGTTGTGCAGTTGATGTCCACATTAATA
TACTGGATTTTCCACGTTAGAAGAATGTTTAAATTTAGTATATGTGGGA

FIG. 4 (49 of 61)

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CAAAGTGGAGACACACAGATTTATACAGCACATACTTTTCTTCATTCA
CTTCTTTGTACTTAAGTTTAGGAATCTTCCCACTTACAGATGGATAAATG
GGTACAATGAAGGGCCAATAGCCCTCCCTGTCTGTATTGAGGGTGTGGGT
CTCTACCTTGGGTGCTGTTCTCTGCCTCGGGAGCTCTCTGTCAATTGCAG
GAGCCTCTGAGGAGAAAATTGACCTTTCTTGGCTGGGGCAGAGAACATAC
GGTATGCAGGGTTCAGGGCTCCTGACGGAGTTGGGGCAACCCTGGAGATAA
GCTCACACAACCCCTGCAAGACCAGGTGCTGTTACCCTAGCCAATCTCATG
GATGAACCAGATCAATGCCAGATGAGCTCTGCCTAAAAATGATTTTTTGGT
GAACTCTGAAAAGTGGAAATATTGTTTCTGTAAGAATATCCATCTGAGACT
CTATCTCTTGGTAATACCAAGAGTTATCAGTTTCTCTTTAACCGAGACAC
CAGCAAAGTGCCTGCTCCAGGGTACTGCCCAGGGGAGCCCTCCATTTGTA
GAATGAATGAGAGTCCAGGTTATGAACAGTGCCTGGAGTGTAGGAACACC
CTCCTTTGCCTCTTTGACAGGTCTGCATCATAACACTTTTTTTTTTTTT
TGAGACAGAGTCTCACTCTGTGCGCCAGGCTGGAGTGCAGTGGCAGGATC
TCGGCCCCCTGCAAGTTCGGCCTCCCGGGTTCACACCATTCTCCTGCCTC
AGCCTCCCCAGCAGCTGGGACTACAGGCACCTGCCGCCACGGCCGGCTAA
TTTTTTGTATTTTAGTAGAGACAGGGTTTACCATTGTTAGCCAGGATGG
TCTCGATCTCCTGACCTTGTGATCTGCCCGCCTCGGCCTCCAAAGTGTT
GGGATTACAGGCGTGAGCCACCGTGTCCAGCCTGTAACACTTCTTATAGC
ACTGAGTTGAAACCTTGCTCCTCCTGGTTCCTCCAGGAACTGAAATCTT
TTTGAGCCAAGTCTAGCACAGTGCCTGGCATGTACATTCAAGTGGTAGAG
TTTGCTGCTTGAATGGGTGAATGGGAATTTGACAGCATTTTTATTCAAAT
TAGTATGTGCCAGGTATCGTGCTCGCTCTGCATTATCCAAGGGAGTGAGC
CTCTGTGCAAGTATTTGAGACACGAGGGAAATAGGTTCTACTGTGGGAAA
AAGAGCATTTCATGGACTTGCTCTCCAAGCAGCCTTCTGATTTTTAATTT
GGCTCCCAGTATCTTGATATCAGGAGTCAGTCACAAGAACTCCATCTTTA
GTAAGTTATATTTTCCACAGGAAATCTAAAAGCTGTTCAACATGTTAGTT
TCCTGTGAATTTGATAAGCCATAATCCATTCCTAACACTGAGCCCTCCTG
AAATTTGGTGTCTGGTCTGCGATAGCTAAAAGCCCTGTCTGGGTGGCC
TAGGGGACTCCTCTGTTTTGCCTCCACAGGATCCACTTTGCAAATTAACC
ACTGGTTCTCCGTTGTAGGAAGTCCACCTTCTCAGAGCCTGTCTTTC
TTCCTTCTTCTCCTTCTTCTCTTCTTCTTCTTCTTCTTCTTCTTCTT
TCTTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTT
TCTTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTT
TTTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCT
TTTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCTTCT
TTTGTCTCTCCCTCCCTCTCTCTCTTCTTCTTCTTCTTCTTCTTCTTCT
CCTAGACAGGATCTACCTTTATCCCCAGGCTGGAGTGCAGTGGTACAAT
CATGCATTCAATGCATGATCACAGCAGCCTCAAACCCCTTCTCAGAGTCT
TTATGCGGCAACCAGCAGGGTCTGGAGGGTTGGTGGCTCTGTGAAGTCTC
CTGACAGAACACAGAGATGTCTTTGGTCTGTTGATGTGATTACAAGCTGA
ACGAAGGAGGATCAAAGCCAGTGACAGGAAGGGAGATATGCAAGGGACCC
GAGCATCAGCTCTGAGTTAGTCCATTCTGCTTCTGGGACTTGGGATACAG
GTCAGAAACCTTGAGCTTCTACTTCTCCATCTTCCAATTGTAGCATCCAG
GACCTCAGAATCTGCCAGCTAAGAGGAGCCCTAATGATTGTCTGGTGGGA
TATGGTGGGACCACAGAGATGAAGACATGAATAGCTATTTGAATGTGAAC
AGCAGACGAAGAAATCAAGGCTAGGAGGGTGAAGTGAAGTCAATCAATAG
CACAGTGTGGTTGAAGCAGCACTAGTATCCAGGTTGCATGAGCCCCTGAT
GCTTTCTGCTCGAGGGAAATTTTGAGCCATGGGGCAATGCCCCCTGACGT
AACAGTCTCCACAGTTCTGCCATGTCTCATCCTGGCCCTGTAACCTGGAC
CCAAATCTGCTACCATCCCATCCATCTCAGGAAGTGAAACCTCTTATGTC
AAATAGGTTGTGCAACGTATGTATCAGATCCTGTCTTCCAAGGAGACCG
CTCAGGCCACAGCACTTCTTCCGATCCCCAATGAGCAGAAAATATCTCG
CTATAAACATAGTTGGCACTAAGGAGGGAGTGAAGAGTGATGATGATG
TAGATGGTGTATGTAGCCCCAAGGAAGTGAACAAGCAGAGATGGGGAGCT
GGAAATGCCAGGATGCTCCAGCTTTTGGGGAATTATTAGCTCTTGAGTC
ACTAAAGCCTTCTCAGCTGCAAGTTCCTCTTTACCCTGTGAGGTCTTTC
TTCCAAGACAGGAGACTGACATTTATTCAAAGCAGCAAGTGCCCTGATAC
CATCTTGTGTCTAATCATGGGCTTCGAGCCAGTTATCAAGGTTGATCTC
ATCTCATTTGGTCTTCAATCATTTTGAACAAGAAGACAAGCAAAATAATCA

FIG. 4 (50 of 61)

TGGGTTAGTTCTTATATTATTGTGTGTACATGCAGTGATGTCTGTTCTTT
GTAGTGAGCTGTTCCTTCCTTGTTCCACCTCTTGCTTAGAACAGAACTAA
CAATCTSCCCCCAACATTTTCCCCAATTTCCCATCTCATTCTTGGCACT
GGCTGCTTAATTTTGTCTTATGAGTCATTTTCTTGATCATTTCATG
AGTCCCTCTGGGATCTTAAAGTATGAAAAATGTTGTGTGTACCCACACCT
GTCTTTGTGGATATTTCTCTCCTTTCCCTTCTGCTTCTGGGATTATTTGG
GAATGGGCACTATGATTTTTATCATATCGCTTCCACTTCCCTTATGGCAT
CATCTCCAATGGGCTTCTTCTCCCTCTTGGATCCAGGTTCTCAGATTGGG
GACATGCAGAGTCCAAGGAACATTCCATTCTCCTCCCTGGTCTAGAACAA
GGAGGCTTACATATATGAGCAGGTGGCTGGGGCTGGCGAGCTATGTAGT
CTCCAATGGCTTTTCCCTGATGTGCGAGTTGTTATGTCAAGTTCTGGGAGA
CCAATAAGACCTTGTCTTCTTGGATCCATCAGAAAAAGCCCTGGGT
GGGTAAGATGGATGGCAGGGCTCTCCTACTCTATGTCTTTTCTCACACCT
AGTGGGTATAAGAGAGGGGACCACAAACAGAGGGGGCTCTGGTACCACTT
ATCCAGGGTCTGGAACATTTTCTGTAAAGGGCCAGATAATAAATGTTTC
AGGTACAACCTACTCAACCTTGCATCATTTCAGAAAAGCAGTCAGATAATA
CATAAATGAATGGGTGTGGCTGGACTTGTCTGCGGTCCCTGTCTTATA
TCATTGTATATATCATTTTTTCTTACATACAAATTTAGAAGCAATACTT
AAAAAAAAAAAAAGCCGTCTTATTGAGCACCTACTAAGTGCCAGGTACCT
TTTTTCCCTCATTATCTTATTAACCTTTCATAATAACCTTTAAAGTAGA
TAATATTGAACCATTTGACCTATGCAGAACTGAGGTTGAGACAATAAAT
TATTTAAGACCGCACAAACAGTAAATGCTGGAACCTACGACTCAAATATGG
GTTAACTGAACCAAAACAGATCTTTATTTCTCACTTTTAATTGTTACAT
ATGTTTATTGCTCATCTCCTGTCCACATGGTGCCCATCGGCAGACTCCT
TTCTCATTCTCAGTGATTGAGTGACATTCTAACTACATTGGCCTGGCAG
ATTACCTCTGTCCCTTAAATGTTTCCACATTGTCTTTTAGGATTGAGA
TCCTCTCTGTTCCCTTGTCTTCCCTCCTTTCTTCTTGGCGGTGACGTG
CTGTGTGAATTTGTTTCTTTCTCCTCTCAGGGTAGTACTGGGACTTTCCA
AATCAGGGTTTTAATGATCTCTCTTCTTCTTCTGAATTTCTTCTTAT
TCCCATTCACTTTCTCATCTATAAGTGGCANCTTTGTTGCTGGAAGATAT
CCCTTGTGAGGATTNCTCTTTAANAATTTGTCTNNNACC

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GTGATCGTCAACCTCCCACCCTGTAGGGCCTCAAGCATTGAGGACAATCA
CTGGCTGCCCATTAACCCAGAAATGTTGCCGAGACAGGAGGCCGTGGCCC
AAGTTCTGGAATGGGGTATTATTATGTGAGCACAAAGGCCTTTGCACAA
ATGAAGGCTTTAAAAATGCAGTCTTAGTCAGGTGGAGGAGGGCTTATAGG
ATTCCCAGGAATCTGGATCATTCTCTTGAGAGCTTTCCCTTGTCTCTGTT
AAAACTCACATCGTACGGCCCAAATAACAACAAAAATGGATGTAAATTC
TTGAATAAATCTGTGGATGGGGGAACAAGGCCACCCCCAGATCTGCCA
GAAGCTTCAGGTGAGGGTCCCAAATGCCAAAAAGTCTGGTATCAGAGAGG
ATGGCCAGTGACNTGGGGACACATGCCCTTTGCTGTGTCACTCAAGGAGC
AGCAGCTTCGGCCCCGCACAGTGACCAGGACCCTGGCTTCCCACGCTGGG
CAGGAGCTGGTGTCTGATGAAGGGAATGCCTGGCAGCACGTGCTGTCTGT
CTCCTCGTGTGACCTTACCTGGCTTTGCTGCCAAGAGGCCACTTGCAATTT
CTTTATTTTTTATATTTTTTTAATTTTTTTAAATTTTTTATTTTTTTA
TTTTTATTTATTTATTTATTTTTTAAATTTTTTTTTTAAATTTATG
CTTTAAGTTTTAGGGTACATGTGCACATTGTGCAGGTTAGTTACATACGC
ATACATGCCCATGCTGGTGGCTGCACCCCACTAATCGTCATCTAGCAT
TAGGTATATCTCCCAGGTTAATCCCTCCCCCCTCCCCCACCACCAAC
AGTCCCCAGAATGTGATGTTCCCTTCTGTGTCCATGTGATCTCATTGA
ATTTCTTTAAAGGTGGAATCTCTCAGTGGGTCTAATCTGTTTCAAGAAAT
TCAAAAGAGTATCCTTGGGAATGACTGGAATCCAGAGTCATCTGGTAAT
CCTCATAAAACAACTCCTGGATGTCTCTCAGCACATCTCCACCTTGAAC
GCAGGAGGCTGGTTCAAATGGAGGAGCATCGCTCTACTGCACTTTTTTTT
TTTTTTGGCCTAAAGTGCAAAAGGGGATACGTTTCATGTAAATAAATCAA
CTGCAAAATCGTAGTTATGCTGAGCCCTGTCCCGTGTGTGGACACAAAG
GAACCAAGGCTTTTCTCCCGCCCAACACACACATAACACACACACAAA
ATCATAAAAAACATACATACCCCAACACATAACACACACACACACA
CAAAATATATACACACAACACACACCAAAACATGCCCAACAACTGTGTCC
AAAAATAAATCCTACTGGTGGGTTTGTGGTCTCCCTAACTTCAAAATGA

FIG. 4 (51 of 61)

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AGCCGTGGACCTTCGCACTGAGTGTACAGCTCTTAAAGATGGCATGGAT
CCAAAGAGTGAGCAGTAGCAACGTTTACTGTGAAGAGCAAAAGGACAAAG
CTTCCACAACCCAGAAGGGGACCCAGCAGGGTTGCTGGTTGGGGTGGCC
AGCTTTTACTTCCTTTTGGCCCCCTCCCATGTTCTGTTTCCATCCTATCAG
AGTGCCCTTTTTCATCCTCCCTGTGATTGGCTACTTTTAGAATCCTGC
TGATTGGTGCATTTTACAGAGTGCTGATTGGTGCCTTTTACAATCCCTT
GTAAGACAGAAAAGTTCCCTGATTGGTGTGTTTACAATCCTCTTGTAAGA
CAGAAAAGTTCCCCAAGTCCCCACTGGACCCAGGAAGTCCACCTGGCCTC
ACCTTTCAACTCCATAATGGCATGAAAATACATATGTTGTACAAAACATA
CATAACAAAGTATACATGCATCTCCCCAAATATACACATACCACAGAAA
CATAACACAGGAAGTCAAGTACCTGTCAAAAGTCTGCATGGTATTGCC
TCTGCAGTGAGTAGTTAGAAAAGTGAATTTGTTTTTCAATAAATTGGAGT
CCTTAAAAATCGTTGTAAGATAGAAAATTTTAAAGTATATAAAATAAA
ATATGTATGTCTTTGGTCTAGCATTACACATGTAGGAATTTATCCTAG
TGGAGTAATCAATGATATATGCAAAGATTTGGACAAGCATATTAAGCACA
GAATTATGTATGCATATGTGTGTATATATATATATATCTGATACATAT
AATAATGTAAAAGTGAATAAAGTCAAGATGTTCAAAATTGAGGATTAGTT
AGACTATGATCTGTCCATATGTGACATACAAGTTAGCTGCCCCCTTATTCT
CTCGAGCTTCAACCTCCTATAAACAGTGTCCCTTGTATATCAGTATTGGT
ACAGATAATCGAAGTTATTGAGGTTTTACATGGGGCAATAAAGGCAAGAG
TTTATGAATACTCCATACTACACTAGGTAGCACCCCCCTATTAAAGACAAA
CTCTTCTCTCTCATTTCCCTTCTTTCCGGAACCACTGGTTGAATCTCT
ACAAGTCTCTATTGCAAGTGCCTCAACATGGCACCCCTCCCTGCATCTCCA
TCTTCCCTGTCTGAGAGCAATGGCCTGTGCCCCACACTCACATCCTC
ATTCATTCCAGAAGTGAGCACCACAGAAAGTGCCTACAGTTACCCCAACCA
CCTTCTTAGAAGATAAGTTAGTGTGTTTTGACTTTTAAAAATTTTAC
TTCTCTTTTTCTTCAATCTCATCCCATCCCAAGAGGTTTATCAAGAA
GTTCTCTAAAGATATGTGTCTCCTTATGGAATTTAACAGAAATCAGGGAT
TTGTATTCTAGCCATCAAGGGAATAACATTTTCCAGGTCTTTAGACAAA
TAATGGAATACCTTGCAAGTAAATTAGATACACTATTGTAGAAAAGTATTGA
TGAAATGGAACGATGTTTGAAGATATCATATTGAGTAGAAAAGGCAAGATA
CATTAAAGTAGGAAATGTATCTTACAAAATAATTTGTCAGACACACTCCTA
TATTTGTATGTTATATAAATGCGTATGTGAAGAAAGGCTAGAGGATGAGA
CCACAGTCTTCGGTGAAGTTTAAAGAGATGAGGCTGCAGCATGCTCAGAAA
GGCCTGGGTTATAGTTCTTCCAGTAATTAAGGATGTGATCTTGGGTAAAT
TGTCCATCCTCTCTAAACTGCACCACCTTTGTCTGTAAACAGGAAGGA
TGGTATTTACCCCAGGGTCATCAAAGGATTTGGTTGGAGAAAAATAAAT
AAATGGGCTGAGCCCAGACCTGGCACAGTGAGAGCACAGTGGTTGACTAT
TGTGCTGGCCTGTTGTTTCTGTGTTATTGACATGCTGCTGGTGGTGGTCC
AGAAGCTATTACCTTAATTGGTTATGTGGATTCCCCTCATACTGAGCAG
CTGTGTGTGGTGTGTAACCATAGCCATACACAGTAAGTACAAAGGGCA
AATGTGATGGAAGAAATGCAAGGAAGTGCAGATAAATAGCTAATGGGCTGT
AGAAGGAAGCTAGTCTTGGAGGGCTTGATCAAGGAAGGTCTTTTGCAT
GTCACCTTTGAAGAAGAGGGGACATAGAAGAGGTATAGTGATCCCGGAG
TGTAACCTGGAAGGGAACATGAAAAGAGGACATTTTCTCTGGGACATGGG
GACTCCACTTGCAATGAACTCTGGAATTGGGGCAAAGAACCATCATGAGAA
CAAGGGCTTCCCTGAACCTCCAGGCTCATTGGCTGATCTAAACCTGTG
TCCCCTCTTCTTCACTCTCCTCTGTTTTCTATACCTGTATTATTGGAC
TGGACTGGAAGCCACCTGATCTATCACAAGTACCTGAAATGTGTTGAAT
AGGTGTGGCACAGTCTTAGCAGAGTGGCACTACCCACAGGAATTTGT
TTATACCTTTGGCATGGAATAAGCAGGAATGAGTGATCACTGATAACT
GAGGATGCTATTTATTATTGGCCAAAGGAATACTTGTGTTGATTTGCAT
AACCCTCACAAGTCTGTTGATTACAAATGAGTACCAGACCTAGCTCCTTC
AAGTAAAGGATCCTGAGAACTGAAGGCAAACAGAGCTCCAGGAGTCCAAG
ACAGAGCCACAGACCACGAGGATCCCTGGCCAGGTAGGTGGTCTCCTG
CACTGGCTTTCAAGGCCAACAGGATGGATGGGGAAGTAGAGTAGCATCTG
GCCATCTAGACCTTGTCTTTTATCCCCACTGGAAGCACATCTGAATTC
TAAATATGATCTCTGAGACCTGCCAGAACACCTTGTCTCAGCCCCAGT
AGCAGCCTGCTCTCTCCAGGAGGGCTTCACTAACAAAGTAGGGCATTGC
TGGAGGGCCAGGCAGACACTAGCTTAGGAAATCCACCAACCCTGGAAATG

FIG. 4 (52 of 61)

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CTAGTCCCTTCTCTGAAGGCTCAGAAGACTGACTTTAGAGTCTAGAAAAT
 ATTGGTCTTGGGAACAGATTTTGGAGTGCAAAGAGATGGACTTCAGATGG
 CCAGATGCAGTCTCTTTAGGGAATTCTGTGAAAGCTCCCTGCATTTAT
 CTTAATACAGGCAGCAGATTTCTGAGTACCCCGAGGGATGGCCCCAGG
 TCCTCCAGCCTGTGAGCATCCTTCTGTCTTTCAGCAGCACCACAGTATCT
 TTATATGTCTTTGGATACCTACGTTTCTGCCAGACATCTCTTGCTCTGAT
 GTCTGGCTGCCAAATTCTCTGTCAAGCGCTCCAATTTTTGTGTCTT
 TGATTTACCCCAACATGACAAAGGCAGTTGTGCTTCATGTATTCAGGGAT
 ACTGCCAAACCACAAACAGGTTAAAAATCAAATAGCAGATATCCCTGTTCC
 TAAAGACCCATCAGCTCTACCCACCTGCTCCTGCTCACCCTCCTTATTGT
 TGAGTCTTGAAGCCCTTCTTGTCTATTTTTATTTTTTGCATGAACAATTT
 AGTTCCCTTGTCTCACTCCTAAACCTTCTCAAAGGATTGGATTTGTAC
 ACAAACCTGCTATCTCTGCAATCTTAGAAGTGATATGATTCTGAACAAAT
 CACTTAACTTTTGATTTTTTATTGGTAAGATGGGAATACCAATTTTTGCT
 CCATTTCTGCTATGTTGGCCTGGGCTGATGTTGAAAGCTCTCGGTCAA
 CTGAGATAGGGTGTGCAAAATTTATATATATAAATATATCTCTCCAACC
 CCTCCCAATGAAGCAAGTCACGTGAGTCAATCCTACCCTAAGATATTAGG
 GATTGAGCCTCCTGGGACATTTGGTGGCTTAGGTTTTCATGAAAAGAGGT
 TGCAGAGCAACTGCTTTTTGTTAGGCAAAGATTAGGCTACTGCAGAGACT
 CAGCAAACCTCTATAGAGGTGTGAGATGGTAAGTATTTTAGGCTTTGCT
 TGCCAGATGATCTCTCAACTAGTTAACCATGCTATTGTAGCCTCGAAGCA
 GCCAGAGACAATATGTAACAAGAGCATGGCTGTGTTTCAATAAACTTT
 ATTTAAAAAAGCAGTCAGGGACCGGATTGGCCAAAGGCCATAGTGTGCC
 AGCCCCAAGACTAGAGCAATGCATTTTAACTTTTTTATTTTTTGT
 AAAATGCCAAGATCCACAAAAATGCTATTGCACCCCGTGTGTTAGCACTG
 TGACTCAAGGTTTGGGAAATCTGCTTTGAAGGCGTGATAGACAGGAGAG
 CATGGTCTGGCCCCCTTGGTGCCTTCTGGTTGCAGCGAGCATTTCAAAC
 ACAGAGCAAGGCCAGTGGTCTGTTGAGCACTAGAGACATGCAGCAAGGTG
 TCCTGGGGTGAGAAGATGCCATAACTGGTCCCCCTTCTATCTCCTTAGGT
 CTGGACTTCATTTCAATTTCTGTTGAGTAATAAACTCAACGTTGAAAAT
 GTCTTTGTGGGGGAGAACTCAGGAGTGAAAATGGGCTCTGAGGACTGGG
 AAAAAGATGAACCCCAAGTGCTGCTTAGAAGGTAAGGTTCTGTAGAAATC
 TACCTCAGGGCCAAAGTGTAATTCCTAGAGCAGAACTTTGCTAGGTGCTG
 TGACAGACCCAGTTGTTTCTGCTGACTTGCACAGTAAGTGAGCTTTCA
 AATTTCCCTGGACAAATAACTAGACAAAGAGAAATCTGGAAGAGAAAAGG
 AAGCTTTTCTCAGTGTCAGGCACATCAGGTAGTAGATAAAAGGATCGT
 CCTCAGCTTACAGATTTGGGGCTTTAGCATCCTGTTTGCCAACTGGATGGT
 TGCATATGCTTCAAAATGCACCTCTTCCCTCCCAACATTCCCAAGTGGA
 GAGAAGCTCCGATGAGAAGGAACTCTTAAGGCTGGGCTGAACAAATGA
 CCCAGGCACAGGGCATCTGAGTATTCATGAGGAACACATTTGGGTGTTG
 CCCATGGGGGACAATAGGAGGAGGCTTTTGACCCAAATGATTGTCTACTG
 AGGTGTGACGGGAGAGGCCTGTGACATGCCAGAGGCCAAACCCGTGATCC
 AGTTCATCTCTATTCTATGTTTCTGAAGAGGGAAGCTATGATTTAATGTC
 ATTACTATCATGCTGCTCTAGTATTTCTCAGCACATACAGAAGAGGGA
 ATTAATGGTCTTGTATACCCCTAAATCCTTGGAAAATCCGAATTGCATA
 TGCTAACCTCACTGCGTCTGACTGCAGACCCGGCTGTAAGCCCCCTGGAA
 CCAGGCCCAAGCCTCCCCGCCATGAATTTTGTTCACACAAGTAAGGCCTC
 GGGGTGAGGTGATGGGGGTGGCTGAGGTGCGAGGGTGGGGATGGGGATG
 GAGCCATTGGGTCTCTTACAGGGTGAGAGAATTGTAGAATGGGGACACC
 TAAGGGTGCTGGATGGGGCTGAAGTCTTCTCTTGTGGAAGCAAATCCCA
 TTAGGAGATAACTCTGGGAAAGATGAGCCCGGGGAGGGGCAGGTGATGCT
 CACCTGCTAAGAGGCAAGGCAAGGAAGAGTTTGTGCTTGGGAACCTTC
 CAGGTGCCTCTTCTGACCATAGCCAAAGAGACTGGAGACACAGACCTCCTC
 CCAGCACTGAGGACAAACAGCCATGGGGCCAGTGGGGGTGCAGGGACACC
 CACACCACTAAGGGCTCAGGGCGGCGCCTTCAAGCCCTGAACCTTCTCT
 CATGCTGCCATTTGAACACCACAACACCCTAATAGGAACTGTTAACATT
 GCCACTTTCAGGTGTGGAAACCGAGACAGACAGTGGAGATTCCCTGCCC
 TAGGTGACACAGGTAATAAGTGACAGATGTGGAAATTTAAAGGTACTATA
 ACGTCTGTGCTGCTCAGGCTTAAGGCTCCCATCACCTCCTCTCTC
 AGGACAGAGTCAGGAGGCCTCAGCCTGAGCCCCAGCTCTAGTGCAGGTTT

FIG. 4 (53 of 61)

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ATGTGGGAATACTGAGC FCACTAGTAL-ATGGCAGAGAGGACCAAATGG
GACCAGGTGTGTAAGGGTGCCTGGCAGTTGGGGGAGGCTGCTGTGCT
TCTCCACCGCTGCTGCTGCAGTTACCTTTGATGTTTTAGTTTTGTTGTAG
TTACACCATTTGCTGGCTTTGGATCTGCACTGTGTCCACTCCAGGTGGAAC
CAGGCACACAAGCCTCTCTGTGCGGCCTGTCTGACTTCTCCTTGTGAGG
GCTGGGATCTCCTTCAAATCTGGCGGAAGTGGTTCTCCAAGTCTGGTCCCT
CAAACGTCAGCAGCATCAGCGCCTAGAAGTGTAGGAATACACATTCCCA
GGCCCCACCACAGACCTCCTGCCTCAGAAACTCAGGGCGCTGAGGCTCTA
GGGGCTGCTTTAAACAAGCCTTCCAGGTTATCGTGACGCACCTTGAAAGTC
TGAGAGCTACTGCCCTACAGAAAGTTACTAGTGCCCTAAAGCTGGCGCTG
GCACTGATGTTACTGCTGCTGTTGGAGTACAACCTTCCCTATAGAAAACAA
CTGCCAGCACCTTAAGACCACCTCACACCTTCAAGTGGCCTTGAGAAAGA
TTTGGGGTCAAGGATCATGAGCGAGAACCACTTAAGAGGATAGTGAAC
TAGTCTGCATGTGAGACGCTGAGATCCTATGTCAGGCTGTGATAGGAGGG
AAACAGAAACCAAAGGAAAGAACAGCTTTAAGAAGCGCTTAAGAGGTACA
AAGTAAAATGATGGTGTAGAAAAGTAGCTTCTTAAAAAGAGCATTTTCC
AGTCTCACCCCTGGACTAACTGAATGAGAATCTCAGGAGTGTGAGGCCAG
GTATCCATGGTCTTAAAAATGCCACCCACCAGGTGATCCCAAGTGTGCACC
AGGGGTGAGAGTCACAGCCTTAGGCCATGCCACTCAAAGGGTGTCTTCAG
ACCAGCAGCACCACAGCTCTGGGAGTGCATCAGAAAGACAGAGGCTTGG
CACCACCCACACCTACTGAACCATAGTTTGCAGGTGATTTCTTGCACATT
AAAGTGTGGGAAATGGAAAAGCTTAGAGTTCAGCTAGCTCGGTGACTCTC
AGTCAACCTGCACCTGCTCCATGAACCTCAGACTGCCTGGGATGGGCCAG
AAAAGCTCCTGAGGAGATTCTGATGTAAGGCAGGGCTGATAACCATGGAT
CTCATCTGACCCCATATCACTGGGGAGTTACTTAGGATCTTGCCTGGGGC
CAGTCATCTCTTCATAGACACTGAGAGTGTCCACGATGCTTGGGGCACT
ACAGGGTGGGAGGTGGAGGATCACGGGTGAGTCAGATAGGAAGCCTGCTC
CTGGGGAGCTTACAGTGTCTATAGGGCAGCAAGCCAAGGATGCCAATACCT
GTGTGCAGGTACCACTGACGAGTGCAGAGCGCTGCAGCACCAGAGAGGAA
GCTACCCTGTGCAGAGGGGGCTGAGGAGGGCTGCAGGGAGATGACAGGAA
AGCCGGTGTACAGGAGGAGTCTCCTCCCACTCTTTGGGCATGAGGAGACC
AGGAGGACATTCTACAGTGAGAAACCCAGGCAGAGGCCATGTGCTTATGG
CATGGGAAAAGAATGACACCTTAGACTTATTCTCTACATTAGAATTGCCT
ACCACAGATACCCATATTATAGCTTCACATAGTGTGGTGGTTACTGTGTT
TTCATATTGTACATTTGCCATTTTCCAGCCACCCACCCATTCTTGACAG
TCACTGGCCCCAGCCTGGGGGGCCCTGTTCTTTATCAAACAAGTGCCTGAG
CTCTTTGCAGAGGTGAGGGTCACCTGTCCAATCAGAGGCCAGGAGGGAAC
GTTCCCTTTAAGACCCTACTCTAGGCAGGCCTGGCCCAAATGAGTTGCT
AGGAGCCCACGCCCTAAGAACCCTCTGAGCACTGTTGTGGCTGGTCCTGC
TGCTAGAAAGTGTCTCCTCCAGGGCCAGGTGCAAGATTTGTGGCTTTTCAA
AGGAGCCACTAAAGCTCCAGCTCAGCCTTGCAAGGTGCTGGGCTCCTGGG
GGCTTCTGCTCCCAACCCTCCCAACTCTTCCATCACCGCTCCCTTAGCC
TGGCCAGTGCAGGGATCTGTTCCACTCTAGGCACTGCTGAGGGAATGATG
CCTCCAGTCAGAGGGTGCAAAAAGAGAGTTAAGAAAAACAATGATTATA
AAAAGTCTTTTTATACGCCAGACATTTTCTTGCTCAGGCTAAGTGCTA
CTTATTTGAGTAAGCATTTTAGTTCTCATAACTCCTCTCTCAAGTAGGTG
CTGCTATTACTTTCAATTCACAGATGAGGACATTGAGGTTTGGAGAGACT
TAGTAACTTGTCTCTGTCTACAGCAGAGCTGGGATTTGAATCTATCTG
TCCAAATCTGGAACCCATTGCTTGACAGAAAGCTTAATTGCTTGTCCC
AGCAAGATAGAAAGCCTGGGAGTGGAAAGAAATATTCAAGTGGCTGTGATGT
CTGAGCCCCACAGGCAGGGTGGAGAGCTAGGGCTGGGGCCCTTGGACGTGG
GGAAGAAAAGGGCTGAGTCTTCCATTTTCAATGTGAAGTGTGATATCTGG
TGATATTGATCTAGGTCCAAAGGTGAAGAACTTAAACCCGAAGAAATTCA
GCATTATGACAGGATCACAAAGTACTGGTCTGGACTCTGGGAATCTC
ATAGCAGTTCCAGATAAAAACCTACATACGCCAGGTGACTCTCAGTTTTG
GCTGTGTTTTCTGCCTCCACCTAGCAGGGGTAAGGCCTCCTGCTAGGTGG
GCTCAACTCCATGCTATACCATGCCCCATCTCCAGCAGGTGGTGAAGCG
AGGAGGAGAGGGCCCAAGGACTAGGGCATCAGATGAAGGGTCTCTAGCAA
TGACCAGATCTGAAAGTAGTCTTTCTGGAAGGGCTGGAGAAAAAGAAGGA
GGCAGACACTTAGACTGGAAGAAGAGGAGGCTTAAACCGGTGTGATGGAG

FIG. 4 (54 of 61)

GGAGAAGTGGACCACAGAGTCAAGGGAGAGGGACTGTGCATCAGGCCTGA
AACCCACAGCAGACAGGAGAGACCTTCCCTGCTCTCAGAACCACACATG
TTCTGACTGTCTTTTTCCAGAGATCTTCTTTGCATTAGCCTCATCCTTGA
GCTCAGCCTCTGCGGAGAAAGGAAGTCCGATTCTCCTGGGGGTCTCTAAA
GGGGAGTTTTGTCTCTACTGTGACAAGGATAAAGGACAAAGTCATCCATC
CCTTCAGCTGAAGGTGAGAGTTCTAGCTCAGTTTCTGGGCCTTTGGCTA
CCCAAAGTAAAAGGCCAAGATCCTCAATGCCTCTCGCTTTCCTGCAAAT
TCTTATCTTGGCCAATATAACAGGGACATCCACCTTCTGGAAGCACCAG
GCAGAAGAGCCCCATAACTTCTTCTCTGGTTCTTGGCCCTTCTAGGGAA
GGAGGAGAGACTCCTCACAGCGGGGAGACAGCAAGGAGCTGAGCACCTGT
TCTCCTCTCCTGGGCTCACTGGTCTGGCCCTGGGCGGGTGGCGGTCCCC
TCCTGCTGTGGCCCTCCATGTGGCAAGCAACACAATTGGGCCAGGACCTT
GGCGTGCTGCTGTAGGGTAGGAGGGTGTGAGGGAGCACTCGGAGGGCAGT
GTGTCTGCCCTGCAAATTTAGTCTGGATGGAGCATCCTTCACTTGAGG
GGAGAAATCTTAGGAAGCTGAATTAGATACAGATCTAAGCCATATTCTCT
AATTTTAAAACTATAGAGCTGAGATTTTGGTATCCATCTGACTCTTACG
TCTCTCTCTCTCTCTCTCTCTCTCAGTTTATTTTTAATCTGGGGGACA
AGAAGGCCTGGAAAAGAGGGCATGATTGCTTATCATCCCTTAAATACCAG
TACCAAGGCTGACACGTCACTTTCCCAAGGACCATCTGCCTTCTCTCTT
TTCCTCCTCTCCTGTGTAAAGGCCTGGAGGATGAGCACATGTGCTGTGTT
TTCCTCCCTCTCAAAGCCTGTGCTATCTAATTAATCCCTTTTACCTCACA
GAAGGAGAACTGATGAAGCTGGCTGCCCAAAGGAATCAGCACGCCGGC
CCTTCATCTTTTATAGGGCTCAGGTGGGCTCCTGGAACATGCTGGAGTCG
GCGGCTCACCCCGGATGGTTCACTGCACTCCTGCAATTGTAATGAGCC
TGTTGGGGTGACAGATAAATTTGAGAACAGGAAACACATTGAATTTTCAT
TTCAACCAAGTTTGCAAAGCTGAAATGAGCCCCAGTGAGGTGAGCGATTAG
GAACTGCCCCATTGAACGCCTTCTCGCTAATTTGAACTAATTGTATAA
AAACACCAAACCTGCTCACTAACTTTCTGTCAATTGGGTTTCATTTCTCA
TTCATGCTTTAAGGATTTGTGTTTTTAGGATATAGCAAGAAGCTTGTTTA
ATTACAAAGTTCTGGGTTGGAAAGAGACCGGCTTCTGCTTGTGTACTGCT
ACCCTGAACCATCAGACATGCATGTGTGTGTATGCTATGATGTGGCC
AGTCTGAGTGCAATACTTGACGCGGAAGGAGCAGCTGGGTGCATGCTGT
GCTCTAGAATTAGTCTTCTACTGGGTTTGGTAGATTCTGAGGGCATT
GATCCTGGGGCAGAAGTGGCTGAGTCTGTGTCTAGGGTACAGTGTGCAAG
AAAGAAATGTAACAGCAAGTCACAATCCAGCCAAGTGATAGTGAAAAGG
GGTAGTTAGGTCCCAGATAAGGAGCAGGGTGACTTGACCTGTGGGAAAGG
CACAGAGACAAGGAATCTGGGTCAGATGACAGCCAGGAGACCAGGTGAGG
GAGGAGCCAGGTACTGTCTGGGAGGCTTGTCAACAAGGGCATGGTCTTAT
CACTAAGCAGGGCTCAGATCCTCATAATGGGGAGTGGAAGGCTGGCCGA
ACAGAAATCAGGGCTGGAAACAGAGTGAGGGGTGGAGACAGGAGACTG
AGGCTTGGAAATTAGTTTTATTAGTTTTAGCTCTTCAGTTACAAGCAATAA
TAATAGCTTCTAGCTTATTTAAGCAACAAGTATACTACAAAAGGAGCTTT
CTAGAAGGATATTGGGTATATTCACTTCTTACTGCTGCTGTAAACAAATTA
CCACCAACTTAGTGGTTTTAAACAATGCAATGTATTATCTTGCACTTATGG
AGGTGAGTCTGGAATGTGTCTCACTGGGCCAAAATCAAAGTATCAGCAGG
ATAGCATTGCTTTGGGAGGCTCTAGGGGAGAGTCAATTTCTTGCCTTTT
CCAGCTTCCAGAGGCCAAGTGCATTCTTGGCTAGTGGCCCACTCCCATC
TTCGCTGCTTGGGTTTTTCTCACTGCTTTGCTCTGACCTCCTGCCTT
CCTCTTTCACATATAAGAACGCTTGCAATTTACATCGGGCTCACGTCAAT
ATCCAGGATACTCTCCGTCTCAAAGAGGCTTAACCTTAAATCACAGATGC
AAAGTCCCTTTTGCTATGTGATGTAACATATACACAGGGTCTGGGGATTA
GAATGTGGACATTTTGGGGTGCCATTATTCTGCCTATCATGTGAAGTAA
CTTTCAAATGGAAGACATGCTGAAGAAAAGTCAAGGATTTCTGGCAG
GCCAGAAATGACAGAAGGCAGAAAACGTTGGTCCCATCACTCAGATGGGT
AAGAGCCAATCACTGCTTTTTGTGCAAGTGAAGAAAGATTGAGATTCCAAGC
AAAGCATGCAACTGCCCTAGTTTGGGTGATGTGTGCACTCCTTGGTCAGT
GAAGGGCAGCACACCTTGATCAATACTCCCTCCAAGACTGTATCCAACGA
GGCCAGTGATGTTCTCAAAGCAGAGCTAGAGAGCTAATCCAGGAGAGA
GGCGTGTGGGTGGTGGGCAGGAAGACAAAGCTCAGCCGTAAAGGAGTAGT
AGGGACAGCACCTTAGGCATGGAGGCTCAAGTGAGATGATACCCATGGGA

FIG. 4 (55 of 61)

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AAAGCTCTGATAAGGTCAJCTCCTTCTGTTTCTGATCCTGATGGTGATG
TGATCAACACCAGCCCAGTGACAAAAAGTACATAGTATATTTAGTAGAT
GTTTCCACACAGAGAAATGGTAAATATTCAAGGCGAGGAATACTCCAAA
CATCCTACCTTGATCATTACACATTCCGTGCGATGTAATGAGTACTTGCAT
GTATGCCATAAATATGTGAAATATTATGTATCACTATATAAAAAGAAAAA
AAATGTGGCCAGGTGACATCCATATTTTGGAGAGGAAGGCATGTCTTCTT
CATTAATATCACAAAACATATTTTCAACAACAGACACAGCTGTTCAAATTA
GTCTCTGAGCCGGGGCTGTCTCATGGCAGTGAGGACTCTGGTTCCCTTAC
AGACTAGCAGAAAGGAGATGGGGCTTACTGACCATGGCCTTGAGGAGGCT
GAACATGCAGGCCAAATGGAGACACAGACAGCCTGGGCTTGGTCTTGCTC
CATCCCCTTCCAACCTGATGAGATATAGTGAGTCACTATGACGTGGGTCA
CTCATGCTTCTGTGAGGCTCCACCAAGACAGCAAGTGCATCAACACCTT
ACGGAAGCACAAGGCCCTGTTTGTGTTGACTTCATGAAAGGCATGGTTG
TGGTGATCGCATTGAGTAGGCTTTTGGGTGAGAGGTGAAAAACCCCACT
ATCATGCATTGCAGCCCTCTGGTGGAACTGTGCTTCAGGCTCTAAATTT
CAGGCTCTAGACTGACTCCAGGATGAGTATTTGGAAGCTGAASTCAATCT
GTGGTCTCTTCTCCTGTAGAGCAGGAGTCAGCACTTTTCATAGAGTGCCA
GATTCTATATATCCTGCCACATGCTCTGTTGTACAGAACAAAGGAGCC
ATAGACAGCATGGCTGTGTTGGCAAATACAAAAACAGGCAATAAGCTGT
ATTTGGCCTTTAGGCTGCAGTTTGCCAACCCCTGCACTAACACAGAGCTT
AAAGGTGGTGGTGGTGTGCTGGAGCTAGCTTATATCAGCTTGCAATAGCC
AATTGCTAACATCTCTTCCAACTCTGTGCTGTGCTTGTGCTTGTATAG
TTTGAAATTGGCTACCCCATTTAATGCTGCAATCTTTTCTCACCCAGCA
CTACTGACTCCCTTTGCCCTGTCTTATTTTCTCACTCTAACATGCTGT
ATAGTTTTCTTCTTACATTTATGTTTGTGCTTCCACTAGCATGTATGT
CCCACAAGTTCTTTGCTCTGTGATGTATCCCAAGAACCCACTGCAGTGCT
TGGCACTTGTAGGAACTCCATAAGATTTTTATAAATGAAGAAAGGAAGAA
AAAAGAGAGGGAGGGAAAAAGGAAGGCCTTCTATTTAAATGATGGC
CTTCTCCATATTTCTATAGTAATATGACTTCCCTTGCAAAGGGGGATGCA
TTTTGGAAAAATGTATATAAATAAACTCAGGTGGTTTTGAATTTCATTTT
CTAACTGTAATTGTAATCATTGGTCTTTATGTTTAGTGAAAAAGTTTGG
CCCTTATGCCTCACACCTGAGAATCCCAAAGTATTGGTTTGTAGAGCTC
CCATAGAGAACCATAAACTGGGTGGCTTAAAAACAAGAAATGTATCGTC
TCCTGGTTCCAGGAGGCCAAAGTCTGAACTCCAGGTGTTGGTTCACTCTGA
GAGCTCTGAGAGAGAATCTGTTCCAGGCTTCCCTTCAGTTGTGGTAGCT
CCAGGGTCTCTGGCTGGTGGCAGCAAACTCCAGTCTCTGCCCCATCT
TCACATGACTGTCTTCTCTGTGTTTCTGTGTCCAGATTGTCTATAAG
GACAGAGTCATACTGAATTAGGGCTCACTCGAATGACTTCATCTTAAGT
GAACTGTATCTGTAAAGACCTTATTTCCAAGTAAGTCAATTCACAGCT
ACTGGGGGATAGGACCTCAACATATCTTTTGGGGGACATAATTCAACTC
ATAATACCCAACATGATAACTGTTTCATCCCATGAAATTTAATGTCTCTCA
AAAGGTGATCTCAGGGCATTTAATCTGTGACAGAACTCCCATAGGAAAC
ATTCCAACCAGAAGCTCCTTTCACAGCTGGTCACTCCTCCTACCCCATCC
GAGGTCTGGGGCAGGGTGAGGCAGGTGGGGACAAGAAGAAGGCTGTCTC
GGGTGTAGAAAAGAGAAGACCCTTATTCACCCGGCACTCTGTTTCATGAATG
AGCTATCCAGCATAGGATATAATAAATCGCTTTAGGAGTGGTAGACTCCA
AACATTTTTTTGGTCCCAGTTATCCTAATCAATTAACAAACTCTAGAAC
CCATCTTGAAGTGCAGGCATTGGGACATTATGAAACTTACACAGAATTCA
AAAAATTTACAAGGGCTAAATAAAACAGGGTCTGACATCTAATATTTTCTT
CCCACATTTCCATGCACTGTCTGGCTCAACCATCCCCAACCCCTCACTCTC
ATCCTGGTGGACACATGCCTAGTGATGTGATCAGCTGGTTTACAGGGGGC
TGGTGATGGTGGATATACAGCTTTTGGCAATTTCCATGGCATAACTACTC
CAAATATGGCCAATTTCAAACCTACCAACATGAAGGCACAGACACAGAGTT
TGGAAGAGATGTTAGCAATTGGCTATTGCAAGCTGATATAAGCTAGCTCC
AGCACAGCACCACCGCTACCTTTAAGCTCCTTGTGTTAGTGCAAGGGTTG
GCAAACTGCAGCCTAAAGGCCAAATACAGCTTACTGCCTGTTTGTGTAT
TTGCCAACACAGCCATGCTGTCTATGGCCTTCTTGTCTGTAAACACAG
AGCATGTGGCAGGATATATAGAATCTGGCAGTCTTAAATAAGTGCTGACT
CCTGCTCTACAGGAGAACACAGATTGTCTTCAGCTTCCAAACATTCTCT
CTGAGTCAGTCTAGAGCCTGAAATTTAGACTGAAGCACAGTTTCCACCAG

FIG. 4 (56 of 61)

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AGGGCTGCAATGCATGAAAGTTGGGGTTTTACCTCTCACCCAAAAGCCT
ACTCAATTTTTTACTGCAAAAACATGTTATCATCATTATTTTTTACTTAG
CCCACCTTTCTTGGCAATTTTCCATAGGAAAATGCATTCTAAATTTCAA
CTAATCAGGGGACTTGGAGCCTCTGGACACCCCTTGTTCCTTGCCACA
GTCCCTTGCAAGGAGGCTTATCAGAGCGGCTCCATGCAGGGGCTCAGG
ACAGGATCAGATGTCAGTTGCACCAAGGGGGCAGGGACAGATCCTCTCTG
CTEACCATGCAAGGGACTGTTCACTGCACCGTCATGGTCTGGTGATT
TCTGGTCCATAAGGGAATTTTCAATGCATCGGGTGATTGTACATCAGC
ACAACACTGTGAGGAAGGCAGAGTGAGAATTTGTGTGCCATTTTATAGG
TGAGAAAACAGATGCAGAGACATTAAGTAACCTCACCACAGTCATGCGGG
TTTTAAGTGGCAGACTTTAGGTGTGTGACTCCTAGTCCAGAGTTCTTT
GCACTGCCCTGAGGTGCTAAACTCTACTGTGCTTTAAGACTCACTTGG
GGAGCTTCTTAAAGAGAGATTGCACAACCTGAGATTCTTGTTTAACTG
TTTTGGGATGTAGCTCAGGGATCTAGCTGCCTTAAAAAATACTCCCA
AGTAATCTGTATGCAAGCGGTTCTTTTTTGTCCACCTTTGAAGAAACT
GCCTCCTCCCATACATTTTCAATTAGAAAATGGTAACATGTTTTCAGCCT
GAGAGCCATTTCTGGGTGACCGGACGTCCGGCAGCCGCTGTACTAGCTTT
CAGTCTAGGCTTAAACACACATGATAGGAGATGTCTACTCCAGATGATA
TGAGTCTGAACCATGGAAAAATTCATTGTGTGGCACATCTGGTGGGTGT
GCACTGTCCCCAGCAGTGAGGCACCCAGTGAAGACAGCAGCTGGGAGAGG
CTTAGTTACATGCAGTGGGACAGTGTGGGCTAGACTGCTGAGCCCTCTGC
AGTTTACTCTGTGTGAGGCAATGAGGGTGAAAGGCTGATCAGACCCACGT
GCAGACCATACCTCCAGGGAGACAGATATCAGTCAGGACAACCCCAAGT
GTAGCTGGAGAAGCAGTGCCAGGTATGACCGGATGTGTATCCAACCAGG
AAATCTGCATATAAATATAAGAGGAGAAAATGAACAGATGTTGCTCTTAT
ATGTAGATATTTATGAAGAGCATATAATTTTGTGTGTGTTTAAGAA
GTTTATAAGTATGCCTTAAATGTATAGTATATACTGTAGGTATTTTTT
CCATTAGATATTTTGTGTCTTACTTATCCACATTGACATTGTAGCAAC
AGTATAATATAACAACCTCCTCTACAAAAGCAGAAGGAAGTGAAGCTTTG
GAAGGAAGCACCCAGTGAGCTTGCCCTTTCAAGTGGGTGCACTGAGCAG
GAGTCAGTGAGGTTGAGATCCTTTGAGAGGAGGCAATCATTAAACCAGGAA
ATCTGCACTGCATCCTGGCCACACCTAACCTTGGACAATGGTGTGTTGGA
GCGCCTTCCAGCTCTTAAGGCTTGCGATTTCTTTCTCTCACTCTTCAACC
ACGATGATTAAATCTTCTCCTACAGAGTTGGACAATAAAGCCTTGAGTTC
CTGCCTCCCTGGTGTGATCAGGAGCATAGACATGGCCAGGAACATGTA
GGTGTCTTTGAAAGCTGAACAAGTTAGTAAATTTCAAACCTCATTTCACC
CACCAGTAAATGGGAATAATAATAAACCTATTTTACATAGGGTTGACAA
GAGGAGTAAAGAGGGATTCAATGAAAGTTCGTTATTATCATTGTAGTAG
CAGTGTGATAATATCAACTGAAAGTTCATTATCATTATTAGTAGCAGTA
TTGATAACCTCTTTTCTGTGCCTTCTCACTGGTGGGCCCAGGCCATCAG
CAATGCCCAGGGTGTCTGATGATCTCTGCTGCATCGGGCACCAGCTGTGTC
AATGGTGAGAACAGTACAAGGGTGGGCAGGGCAAGGCAGGAAGCACCAG
GAGCAGCAGCTTCAATGGGGTGAAGATGTCAGGAGCTTAGGGACAGTCAGA
GCGGGTGTGCCTCCTCTTGTGGAGCCTTTCTGCTGGGTAGGAAGTGTG
CAGCTGTGGCCATGGATTACCTGAATATGGGTGGAATTAGGCATTACAGC
TGGGTTAGCTGTGCCTAGAAGGAGGAACTCTAACTGAGAACTTGTCCCT
ATTGCCACCTCTGATAGGCAGATGATCCATCCATCAGTGGCTGAGCTGAG
GTGTGCATGGGGATGGGTAAGAGCCACACACAGGGCTGATGACTGAGTC
TATTTAGAACAATAGATGTAAATCTGATAATGTAAATGTGATAGATTA
TTTTGTCAATTAGAAATGGTACCATATAATTATATATACATAAACATG
TATACATATACACACATATACATGTGTGTATAAACACACACAGTATTGTC
CCCTACTCATTCCATAAACCTGATGCCTTTAGCTGGGATTCCCAGCTTTC
ACTCTCCTCTGTCTGTCTGTCTATATCCTCCCATCCTGTAATTCT
GGCTTATATGCCACTTCCCTCCCTAAAGCCCTCCCTCAATCCCTTGCTGGA
AGTGACATTTTCTCTTTGAGCTGCCCCCTGCTTGTGCTTTGGTGAGGTCA
GCTGTATTGCAGTACCTTGTATTGTGGTTGTACATCATCGTATAGAATT
AATTTCTGACACATTCCGTATTTTTCAAAGGGCCTAGTGTGGGGCTTTAA
CAGTAACACGCCACCCAGCCAGTTAATTTTTGTATTTTTGGTGGAGA
CAAGGTTTACCATGTTGGCCGGGCTGGTTTCAAGTCTCTGACTTCAGGT
GATCTGTCTGCCTCAGCCTCCTGGAGTGCTAGGATTGCAGGCATGAGCCA

FIG. 4 (57 of 61)

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CTGCACCCAGCCACCTATCAAAAATTTTAAGTGCCATTTTATTTTTTATT
TTTTGTAGAAATGGACAAGCTGATCGCAAAATTCACATGGAATTGCAGGA
GGTTCCAAATAGCCAAAACAATCTTGAAAAAGAAACAAGTTGGAGGA
TTTACACTTTCCAGTTTCAAGACTTAGCTCTTAGCTACAAAGCTACAGTA
ATCAGAACACTATGGTCTTGGCATAAGTGATGCTGGACAGGTGAGCCCA
AAGTGGGACTTAACCTGTGAAGGTTCTTGGCCTTGCCAGGAAGGAATTC
AAGGGCAAGCCAATGGGACAAGAAAACAGCTTTATTGAAGGGGCAGTATT
ACAGCTCCAGCCCTGTTACAGCTCCAGCCCTGTTACAACCTCTGACTACTC
CTGCACAGAAGGGCTACCCTGTAGGCAGAGAGTAGCAACTCAGGGCAGTT
TTGCAGTCATTTATATCCACTTTTAACACATGCAGATTAAGGGACAATTT
ATGCAGAAATTTCTACGGAATTGGTAATAACTTTTGGGTCTGAGTCTAT
CATGGAAGGGGGGGCGGGGAACCTCCCTGGTGTGCCATGATGACGGTAAAC
TGATATGGCGAAGTGGTGGGTATGTCACATGAAAAGCTCCTTCCACCCCA
GCCCTGTTTCAATTAGTCTCGGTTTGGTCCAGTGTCCAAGTCTGCCTC
CAGAGTCAAGTCCCACCCCTACCTCTTAAGGAGAGATGTAAATACATGG
AATAGAATTGAGAGTCCAGAAATAATCTCATACATCTATGATCAATTGAT
TTTCAGCAAAGGTGCCAAGACCATTCAATGAGGGAAAGAATCATATTTT
TTCAACAAATGGTGTCTGGATAACCACATGTGAAAGAATGCAACTGGGCCC
TTATCTCACCCATATACAGAAATTAACCTCAAAATGGCTCAAACACTTAC
ATGTAAGAGCTAAACTATAATATTCTTAGAAGAAAAACAGGGATATATCT
TTATGACCTTGGATTTGCTGGCTGATTCTTAAATGACACTGAAAGCACA
GCAACAAAAGAAAAAAATAGGTAAATTGGACCTCATCAAAATTTAAAA
CTTTTATGCTGGGTGCACACCTGTAATCCAGCACTTTGGGAGGCTGAGG
CAGGAGGATCTCTTGAGCCCAAGAAGCTGAGGCTACAGTGAGCCGAAT
GTGCCACTGCACTCCAGCCTGGGTGACAGAGCAAGACCTGTCTCGAATA
AATAAATAACAAATATATAATTATAGATCTCTGGATCTTGCCTTCGGAG
ACTGACTCAACTAAGTCTGGGTGGGAGCCAGCCATTTGTATTTTTT
GAAAACCTCTCAAATGATTTTACTGTGCAGCCAAGGTTGAGAATCACTGT
ATCATAGGGTTGGACTCCTAACTGGAAACAGTTTGCACCATCAGGTGTCTG
CAGCATTCTGATAATAGTTAAGCTTTTCTCTAGATTTTCTGATATTAGA
TGAGTCATGTTTACAAGTTTTTACCAAGAGACAACTATCTTTCTGCCCT
TACTTTCTCTCTTATACTATTCTAATCCCAGAACCCTTTGGAACCTCCAC
TGAGAGATGAATCTAGAAAGTGACTCTCTTGGCTACAACAGAGAGTAATG
TTGGCCTGTTTGTGCCAGATCCAGTTGGTGTGGTGGTGGGACAGCACCT
CCCTGAAATCCCCTCCTCTCCCGTCAGATTCACTCCCCCATTTGCATCAC
GTACAATCATCACTATGGGTTTCTATTACCTTGCTAGGGCATTGAGAGT
ACCATATATACCAACTATTAGTTTGGAGCCATGGTTCCCAAAGTGTGGAC
TGAGGGGCACCTCAGCAGCTCAGGAGGTGTCTGGGATATTTAAATATT
CTGAAGAAAAACACAGTGACATCTGTGAGGCCGTGAAACCGTTGGCATT
AAATTGTCTCAACCAATTGCTTAAGAAGCAGAACTGGCCAGGCACGGTG
GCTCACATCTGTAATCCAGCACTTTGGGAGGCCGAGGCGGGCAGATCAC
GAGGTGAGGAGTTCGAGACCAGCCTGACCAACATAGTGAAACCCCGTCTC
TACTAAAAATATAAAATTTAGCCATGCATGGTGGCATGCACCTGTAACCC
CAGCTACTCAGGAGGCTGAGGCAGGAGAATTGCTTGAACCTGGGAAGCGG
AGGTTGTAGTGAGCCAAAATCGTGCCACTGCACTCCAGCTTGGGTGATAG
TGAGACTACATCTCAAAAAAATAAATGAGAGAGAGAGAGAAGCAGA
ACCATCAGGTGTTTCTTTTGGCTTAAAGTACTCTGTGAAGAAATTCCTGG
GACACGAAGGATACCATGAACTGAGAGATTTTGGGAACCTCTGCTTTAGA
AGCTGGAGGTAGCATTCTTGGGCACAGTACTGCCTTGGGATCAGCAAT
CCTTTTGTAGGTGCATTAGGTGTGGCAAGACAGCTCTTAGAGTGGGACC
GGGATGTGCTTGGAGACAGAGGGAAGTATGAGCTGCCCCGATAAAGAC
ATGCCAGCCTGGCAGAGTGTAGTACTCATGTCTGTAATCCTAGTGCTTT
GGGAGGCTGAAGTGGGAGGATGCTTGGAGGCCAGGGGTTTGGATCAGCC
TGGGAAACAACAAGACCTCTACAAAAAAGAAAAAATAAACC
CATGTGGTGGCATGCACCTGTAGTCCCAGCTACCTGGCAGGCTGAGGTAG
GAGGATCACTTGAAGCCAGGAAGGTAAAGGATACATTGAGCCATGACTGTG
CCACTGCACTCTAGCCTGGGTGACAGAAAGAGACTCTGTCTCAGAAATAA
ATTAATAAATAAATAAATAATATAGTGGCCATGACATCCCTAGAAAGACA
AGGTCCTGGGAATAGGTAGAAGCCAAGGGAATGAGAAATGAGAGGGGGC
CCTGGAGCTGGAACCTGGGGGAGCAGGATGGCCTCTGAGAAGTCTCTGATA

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GTGGTGTCACTGATGTGTCTGATGTTTGTAGTTGTAATTATTTGCTGGGCCC
CTGTGATCCCTCATATCTGATAGCTCTTTGCTAGTCAAAGTGTGGTCTGG
GGATCAGCGGCATCAGCATCACTTGAGAACTTGTTAGAGATGCAGAATCT
AGAGCCCCACCCGGGACCCAGAAACAGAGCCTGCATTTTAAACAAGCTCCC
CAGGTGATTCTCACACACACTCGCATTTGAGAAGCACTGGGCTAGTTGAC
AGATTCTCAGGCATGGCTGACATTGAAATATCCAGGGAGCAGGCTTGGCA
TTAGGATGTTTAAAGTCTCCAGGTGTTTCTAAAGCCAGGTTTGAGGAA
TTACTGGGCTGATACAAATGTTTTGTGATGATGCTTTGTGTGTGTGTGTG
TG
TGGGTCACTTGGCACCAACACAGGAAACAATGGAAATATGTGAGCCATGA
CAGAAAGGTCAGGAGATAAAAGAAATTAGTGACATGAGAGGTACTCCTCA
GGTGTAGGAAAGAGGGTAGAGCAAACCAGGTTTTCCACCATATGTTGGA
TAGGGGGTCAAGTAAATTTCTACTTAAAAATTACAAACAGGGGCTGGGCG
CGGTGGCTCATGCCTGTAATCCCGCACTTTGGGAGGCTGAGGAGGGCGGA
TCACAAGGTCAAGAGATTGAGACCATCCTGGCCAACACGGTGAAACCGTG
TCTCCACTAAAAATACAAAAATTAGCTGGGCATGGTGGTGCCTGCTTTA
TTCCCAGCTACTCGGGAGGCTGAGGCAGGAGAATCGCTTGAACCTGGGAG
GTGGAGGTTGCAGTGGGCGGAGATCGCACCCTGCAATCCAGAGCGAGAC
TGTGTCAAAAAAAAAAAAAAGAAATTTCAAACAGGATGACCCTAAG
CCTGCAGGACTTGGAGACATCTAGGTGACTGATACTCAGTCACAAAACAT
AATTGGTCAAGGCCTGATGAAATGCACAGCAGACCTTCAGATGGTATGC
ACTCAAGTGATATCCACAAGTCCACCTAAAGAAATGCTATATTGAGACAT
TTGGCATCAATCTCTATCAAACAAAGATAGTCCAAAGCAATGGGTTCCAA
AAACACTTTCTAAGACAAATTCTCTATTTGCTTTTAAATATCAGTCATCC
CAGCCCTTGAATAGAGGAGCAAATGATACCAGTGGTACCCCTACCACAAT
GCACCAAGGTATTATACTCTCATGCTCCATTTCTCCCTCTGTCTACATC
ACTAATAACTCATTGATTTCTGGTGCAAGCCCTGCTGGGAGAAAAAGTCT
ACTCTGTACCTTGGAGCAAGTTGCTCAGAGTAGGTATCGAGGATAAAAT
TTGGAAAGTTAGAAAAGCTATTAGAAGGAGATCCTAGTAGTTGAAAACAC
AGCCTGGCCAAGTCAATGATGCTATTTCTCTCCCCAGCCTTGCTGTGCC
ATAGCTAAGGAAGACAATTTAGGCTTGGGCTAGAGGATGGGAAAGGGCAA
AATTACTGATGCCACAGCCAGAGAGGTATTCTAGTAATCTGAGGGTGAG
GACCACATACCTGGTTGAGGACGTACAGTGTGACAGCTGTGAGTGGAT
GCCTGGAGTTCTGGCGTGTCTTCTAGCACAATGATACCTGAGACTCTTGC
ATCATTGGGAATAATAAAATGGGAGTGGATAGATATGAAATTATGATGGC
AATAAGCAATCAGCTAATAGCTTCATTGATGGGACAGATTAAAGATGGCT
GCAATCCTTTGGTCCAGGTTTGGGATATAGGCAGCATTTGTATTGGAAT
GCTGATAGTCTGAGGCCATGAAAAGTCCACCTGCAGTAGTGGTAGGAGGA
ACAAGCCTCACTTTCTTCAATGTGTGTGACTGCTGTCTTGATTCCCTGGG
TGGCCAGTTCCATTGCTGTGGTCTTTGGTCCACTTGACTCTGGGGTGGC
TCTGTGATGGCTTGACCAATACAAATGATGTGAAATGATGCTGTCTCATCAT
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TGGAATACTTGTCTTGCAACCCATCCATCATACAGTGAGAAATTCTAAG
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GTGAACCACTTAGGACTGTATACTCCAGCCCCAGTTGAGCAATGTGGAAC
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AAATAATCCCCTAGGCTTTGGGCTGATTTGTTCCAGATTACTGGAACAGA
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ACTTTTAGGGAGGGTTGAAGCATAGTGAGGAAAACAGTAGGGGAAGCTAG
AGGAAAAAATGATGCTTGGTATGTAGTGGTGGGAAGTTTAGCAAACTCG
CCTGATGTAATGTGGGAAATTTGAAGAACTCAGAACGATTTAAGGGCATG
TTTTATAGGTCCTTTAAGAACTTCTAGGCCAGGCGCAGTGGCTCATGTC
TGTAATCCCAGCACTTTGGGAGGCTGAGGTGGGCGGATCACAAGGTCAGG
AGATCGAGACAATCCTGGCTAACATTGTGAAACCCCGTCTCTACTAAAAAC
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ACTAGGGAGGCTGAGGCAGAAGATGGCGTGAACCTGGGATGTGGATCTT
GAAGTGAGCCCAGATTGTGCCACTGCACTCCAGCCTGGGCAACAGAGTGA
GACTCCGTCTCAAACCGAAAAAAGAACTTCTAGGGC

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TTGGTCCCCTGGAAGCCTCACACATGGTACACAAAGGCTGTCTTGAAAAGA
AACGTAAGTGTGTTTTTTGGTTTAAATAAAATTGATTATAAATGGATAATG
CAAAACATTTTAAAGAATTTTACTAGCTTACATTAGCAGATTTGGATCCA
GTGATTGTTACATTCTGGTACTGAGCCCCTGAATTACTTCTTTGAGTAAG
GCATTATACCAAAGCTATTGATAGTTGGGCTTATAGGGTGTATGTTTGAA
GAACTACTAATGTCAAACCAATATTTACGGTTCGACAAGAGGACATCAG
AACTGGTAATCCTTATTACCATGACTGGCTGGACAGAATACTCAATGTAA
TGGGATTTCTGCAAATAAAGACGGGAAGATGTAAAAAAGATGCCTGAA
CATTCAACATTAATGAAAGATTTGAGAAGAAATATGTATACTAACTGCAG
CCTTATCAAGTATATGAAAAACACAAAGTTAAACCAGATAGTAAAGCAT
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TTTCAACAGTTGTCAAATCCCCTACCCAAAATGAGAATTTTTTAACAGAAG
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CATAACCACACGAATGGAAGTGGCCACCCAGGAATCAAGACAACGGTCAC
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CTCAGGTGGACTTTTACAGGCCTCAGACTATCCGCGTTCCAATCCACATG
CTGCCATATCCCAACCCTGGACCAAGCACATCCAGCCGAAGAGCAGTG
TAGGATACTCAGCTACCTCCAGCAGGCTCCACAGGACCCACGTGAGACA
CACGGGTACTGAGCTGCATCGGAATCTTGTCCGTGCACTGTTGTGAATGC
TGCAGGGCTGACTGTGCAGCTCTCCGTGGGAACCTGGTATGGGCCATGAG
AATGTACTGTACAACCACACCTGCCCAGTAGCCAAGTTCCTTCCACCGCT
TTTCACAGATCGGGGTAGTGGCTTCCAGTTTGTACCTATTTTGGAGTTAG
ACCTGAAAAGAAAGCGCTAGCACAGTTTGTGTTGTGATTTGCTACTTTC
ATAGTTAACTTGACCTGGCTCAGACTGACCAGTACTTTTTTTCCGTGAC
AGTCTATAGCAGTTGAAGCTGAGAATGTGCTAGGGGCAAGCGTTTGTCTT
CATATGTCATGAATTCCTCCAGTGTAACAACATTATCTGACCAATAGTAC
ACACACAGACACAAGGTTTAACTGGTACTTGAAAACATACAGTAGGTGTT
AACTCAGTGAAATAACCAGGACTCAAAGTAAGATTATTTTGGTACACCTT
TCTTGTAGTGTCTTATCAGTGAGTTGATTCATTTTCTACATTAATCAGT
GTTTTCTGACCAAGATATTGCTTGGATTTTTCTGAAAGTACAAAAAGCC
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GTATAAGTTTGTTTTTTATTTTCAATCTATGACTTGACTGGTATTAAAGCT
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TCAGCAAACAACTAAAATGATTGTACAGACAATGCTTTATTTTTCTG
TTGGTGTGCTTGTGGGAAAAAGAAAGAGAGATCAGATTGTTACTGTGTC
TGTGTAGAAAGAAGTAGACATAGGAGACTCCATTTTGTCTGTACTAAGA
AAAATTCTTCTGCCTTGAGATGCTGTTAATCTATATAACCTTACCCCCAA
CCCTGTGCTCTCTGAAACATGTGCTGTGTCCACTCAGGGTTAAATGGATT
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CTCGTAAGAGTATCACCACTCCCTAATCTCAAGTACCCAGGGACACAAA
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AAGGTTTCTCCCATGTGATAGTCTGAAATATGGCCTCGTGGGAGGGGAA
AGACCTGACCGTCCCCAGCCGACACCCGTAAAGGGTCTGTGCTGAGGA
GGATTAGTATACGAGGAAGGAACGCCTCTTGTGAGTTGAGACAAGAGGAA
GGCATCTGTCTTCTGCCGTCCCTGGGCAATGGAATGTCTCGGTATAAAA
CCCCGATTTTATGTTCCATCTACTGAGATAGGGGAAAACACCTTAGGGCT
GGAGGTGGGACATGCGGCAGCAATACTGCTCTTTAAGACATTGAGATGTT
TATGTGTATGCATATCTAAAGCACAGCACTTAATTCTTTACCTTGTCTAT
GTTGCAGAGACCTTTGTTACAGTGTATTCTGCTGACCTTCTCTCCACTA
TTATCCTATGACCCTGCCACATCCCCCTCTCCGAGAAACACCCAAGAATG
ATCAATAAATACTAAGGGAACCTCAGAGGCCGGCGGGATCCTCCATATACT
GAACGCTTGTCCCCTGGGCCCCCTTATTTCTTTCTTATACTTGGTCTCT
GTGCTTTTTTCTTTTCAAAGTCTCTCGTTCCACCTAATGAGAAACACCCA
CAGGTGTAAAGGGGCAACCCACCCCTTCAATGTGCTGATTGTGAGCGTGCT
TTAAGGTGAAAAAGCATGAATGTAACTTCCTTAAAAAGGTACAGCATC
CAATTCAAATATTTTTGTCTGATTTTAAATGCTAGTTGATGTAGTGCTAT
TAAAATTTTGTTCACATGGACACAGAGAGGGGAACAACACATACCAGGG
CCTGTTGCGGGGTGGGGATGAGGGGAGGGAACTTAGAGGACAGGTGAACA
GGTGACAGATCACCATGGCCACATATACCTATTTAACAACCTGCAC

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GTTCTGCACACGTATCCCATTTCTTTTTTTTTTTAAGAAATAGAAAAAA
AATAAAATTTTGTTCACTGATTCTTCCATTTTAAACCTGTTTGCAATGTG
GTTTAGGATGCCCTTACTTCAGCAAAGGAGAAGGAATAGGAGGGCCTTAG
AATTTTGGAGGGAAAAAACCTATAACATACATTGTACTGTATCAAACCT
ATTTTACATGAATGACACAAGTATTCTGAATAAAAAATAATTGAACATT
GTTAAGAACAAGGTGTCATGTAATTTATTTTTCATAAATAAAAAATTAT
AGTGGCTTAGACTGAAAGGAACAGAGAATTTAAAAAATTAAAAAGAAGCC
TTAGTATATTTTGTATATAGTTTCCATGTGCCATATTTGCCATAATTGG
ATGAGAATTTTGTACCTCTGGCAGGGTGACCCTATATTTTCANTNTATA
AAGCGTGCATCATACC

MVLKCIIPPGIDSQCAI'GVRVTALGHATQRVSSIXQIIPQI.WECIRKTEAWIIHPIII.I.NIISI.QP'GGI'CSI.SNKCI.SSI.QRSASA
I:KGSPII.I.GVSKQEFCL.YCDKDKGQSIIPSI.QI.KIEI.MKI.AAQKESARRPFIFYRAQVGSWNMLESA.AHFGWHTCTSCNCH
I:PVGIXNXVDI'FI.I.GKAQKRGTYSE

FIG. 5

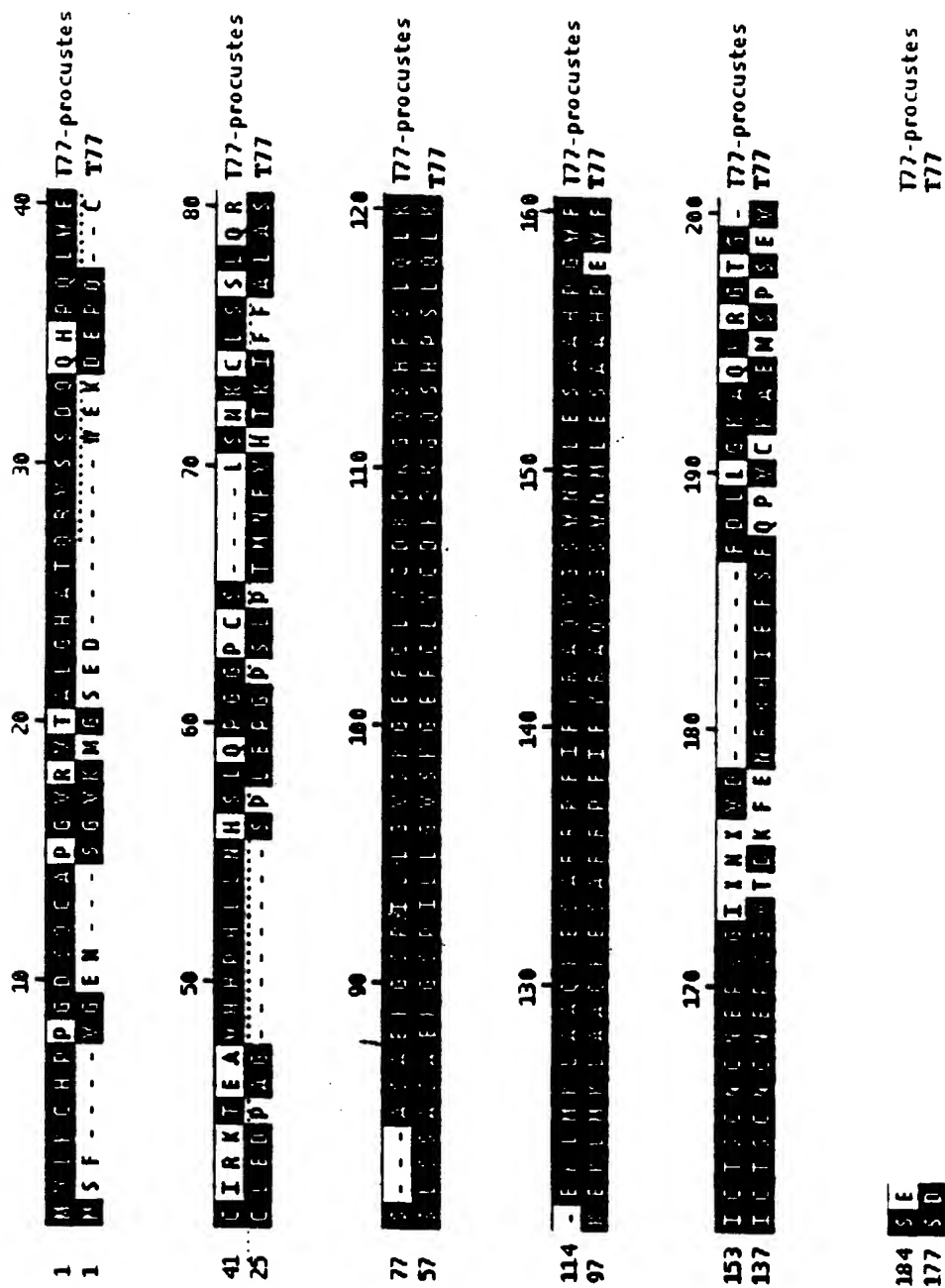


FIG. 6

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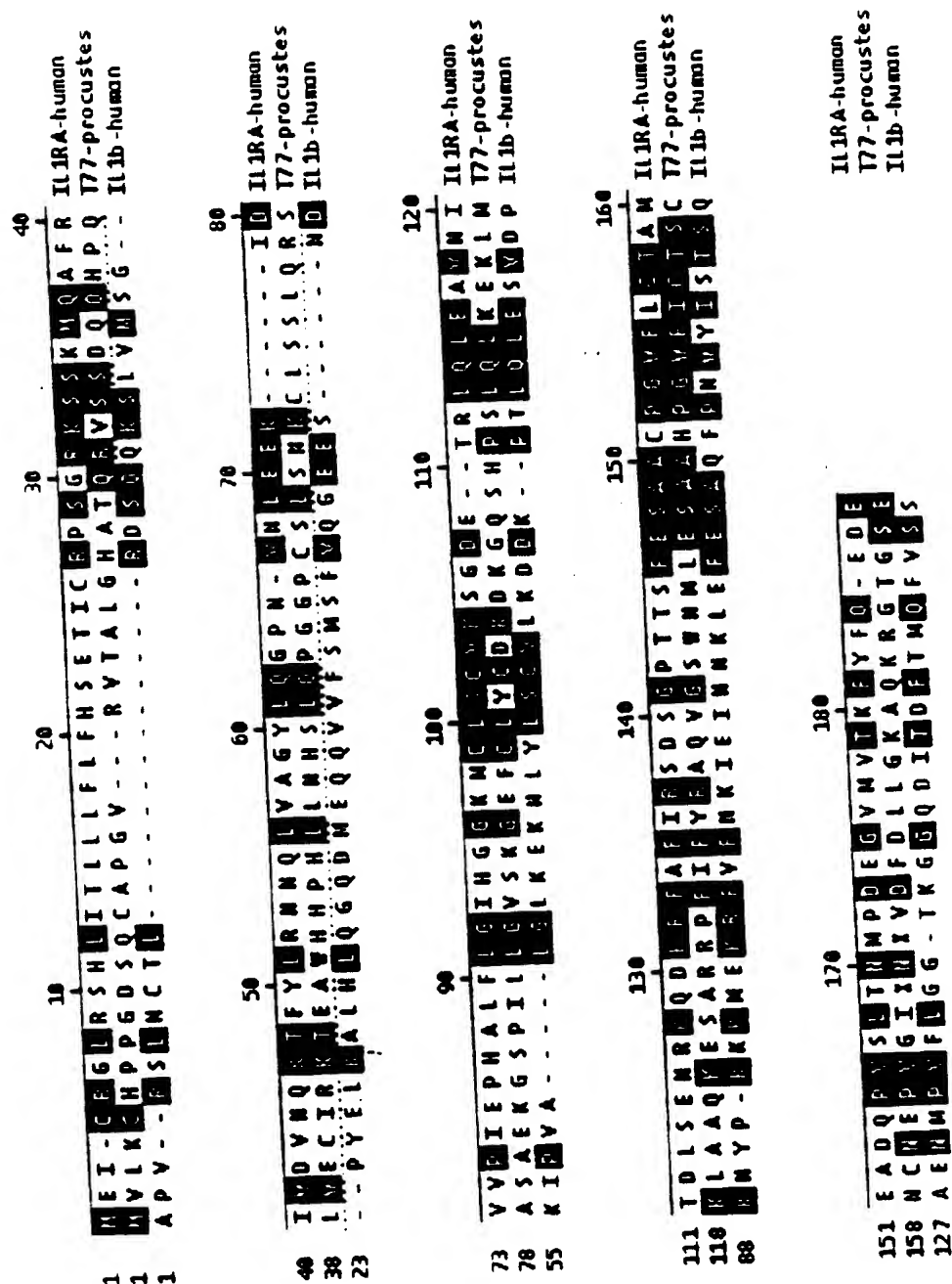


FIG. 7

INTERNATIONAL SEARCH REPORT

International application No.
PCT/US98/16102

A. CLASSIFICATION OF SUBJECT MATTER

IPC(6) : C07H 21/02, 21/04, 1/00, 14/00, 17/00; C12Q 1/68; G01N 33/53
US CL : 536/23.1; 530/350, 387.1; 435/6, 7.1

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

U.S. : 536/23.1; 530/350, 387.1; 435/6, 7.1

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)
DIALOG: MEDLINE, USPATFUL, WPI, BIOSIS. Search terms include author, "TANGO" and protein

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	Database Medline on Dialog, US National Library of Medicine, (Bethesda, MD, USA) AN 09370320. SONNENFELD et al. 'The Drosophila tango gene encodes a bHLH-PAS protein that is orthologous to mammalian Arnt and controls CNS midline and tracheal development'. Development. November 1997, volume 124, number 22, pages 4571-82, Abstract.	1-22

☐ Further documents are listed in the continuation of Box C. ☐ See patent family annex.

Special categories of cited documents:		See patent family annex.	
"A"	document defining the general state of the art which is not considered to be of particular relevance	"T"	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
"B"	earlier document published on or after the international filing date	"X"	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
"L"	document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified)	"Y"	document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art
"O"	document referring to an oral disclosure, use, exhibition or other means	"A"	document member of the same patent family
"P"	document published prior to the international filing date but later than the priority date claimed		

Date of the actual completion of the international search

21 OCTOBER 1998

Date of mailing of the international search report

30 OCT 1998

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